=> fil reg FILE 'REGISTRY' ENTERED AT 15:58:19 ON 07 OCT 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 5 OCT 2003 HIGHEST RN 599148-37-5 DICTIONARY FILE UPDATES: 5 OCT 2003 HIGHEST RN 599148-37-5

TSCA INFORMATION NOW CURRENT THROUGH JULY 14, 2003

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

=> d 174 ide can tot

CN

CN

Chitosan SK 10

. Chitosan VL

L74 ANSWER 1 OF 2 REGISTRY COPYRIGHT 2003 ACS on STN 9012-76-4 REGISTRY Chitosan (8CI, 9CI) (CA INDEX NAME) OTHER NAMES: CN 100D-VL CN Amidan CN BC 10 CN BC 10 (polysaccharide) CN Biopolymer L 112 CN Chicol CN Chitan, N-acetyl-CN Chitin, N-deacetyl-CN Chitoclear CN Chitofos CN Chitolaze CN Chitopearl 3510 CN Chitopearl BC 3000 CNChitopearl BCW 2500 CNChitopearl BCW 3000 CNChitopearl BCW 3500 CNChitopearl BCW 3505 CNChitopearl BCW 3507 CNChitopearl K 20 CNChitosan 500 CNChitosan CLH CN Chitosan EL CN Chitosan F CN Chitosan FL CN Chitosan H CN Chitosan LL CN Chitosan LL 80 CN Chitosan LLWP CNChitosan M CN Chitosan MP CNChitosan PSH

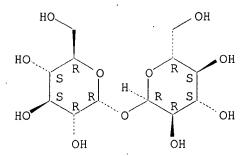
Jan Delaval
Reference Librarian
Biotechnology & Chemical Library
CM1 1E07 – 703-308-4498
jan.delaval@uspto.gov

-1.41.01

```
Chitosol
CN
CN
    Chitosom
    Crystan LA-S
CN
    CTA 1 Lactic Acid
CN
CN
    CTA 4
     DAC 50
CN
     DAC 70
CN
     Daichitosan 100DVL
CN
     Daichitosan DVL
CN
     Daichitosan P-VL
CN
     Daichitosan VL
CN
     Daichitosan VLA
CN
CN
     Daichitosan W 10
     Deacetylchitin
CN
     FCM 117
CN
     Flonac C
CN
     Flonac H
CN
ADDITIONAL NAMES NOT AVAILABLE IN THIS FORMAT - Use FCN, FIDE, or ALL for
     DISPLAY
     57285-05-9
DR
MF
     Unspecified
CI
     PMS, COM, MAN
PCT Manual registration, Polyother, Polyother only
     STN Files: ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BIOBUSINESS, BIOSIS,
LC
       BIOTECHNO, CA, CABA, CANCERLIT, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS,
       CHEMLIST, CIN, CSCHEM, CSNB, DDFU, DIOGENES, DRUGU, EMBASE, IFICDB,
       IFIPAT, IFIUDB, IPA, MEDLINE, NAPRALERT, PHAR, PIRA, PROMT, RTECS*,
       TOXCENTER, TULSA, USAN, USPAT2, USPATFULL, VTB
         (*File contains numerically searchable property data)
                     NDSL**, TSCA**, WHO
     Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
           12052 REFERENCES IN FILE CA (1907 TO DATE)
            2209 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
           12090 REFERENCES IN FILE CAPLUS (1907 TO DATE)
REFERENCE
            1: 139:235940
REFERENCE
            2:
                139:235498
                139:235488
REFERENCE
            3:
                139:235399
REFERENCE
            4:
                139:235340
REFERENCE
            5:
REFERENCE
                139:235291
            6:
REFERENCE
            7:
                139:235290
REFERENCE
            8:
                139:235280
REFERENCE
            9:
                139:235226
REFERENCE 10:
                139:235206
L74 ANSWER 2 OF 2 REGISTRY COPYRIGHT 2003 ACS on STN
     99-20-7 REGISTRY
     .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Trehalose (8CI)
OTHER NAMES:
```

```
CN
     .alpha.,.alpha.'-D-Trehalose
CN
     .alpha.,.alpha.-Trehalose
     .alpha.-D-Trehalose
CN
CN
     .alpha.-Trehalose
CN
     D-(+)-Trehalose
CN
     D-Trehalose
CN
     Ergot sugar
CN
     Mycose
     Natural trehalose
CN
     NSC 2093
CN
CN
     Treha
CN
     Trehaose
FS
     STEREOSEARCH
DR
     229966-89-6
MF
     C12 H22 O11
CI
     COM
                  ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS,
LC
     STN Files:
       BIOTECHNO, CA, CABA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CBNB, CEN,
       CHEMCATS, CHEMLIST, CIN, CSCHEM, DDFU, DETHERM*, DRUGU, EMBASE, GMELIN*,
       HODOC*, IFICDB, IFIUDB, IPA, MEDLINE, MRCK*, NAPRALERT, PIRA, PROMT,
       SPECINFO, TOXCENTER, TULSA, USPAT2, USPATFULL
         (*File contains numerically searchable property data)
                      DSL**, EINECS**, TSCA**
         (**Enter CHEMLIST File for up-to-date regulatory information)
```

Absolute stereochemistry. Rotation (+).



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

5985 REFERENCES IN FILE CA (1907 TO DATE) 297 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA 5999 REFERENCES IN FILE CAPLUS (1907 TO DATE) 64 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 139:235429 1: 139:235194 REFERENCE 2: REFERENCE 3: 139:235002 REFERENCE 4: 139:235001 139:229740 REFERENCE 5: REFERENCE 139:229644 6: REFERENCE 7: 139:229549 REFERENCE 139:229428

8:

REFERENCE 9: 139:227007 REFERENCE 10: 139:226862 => d his (FILE 'HOME' ENTERED AT 15:17:49 ON 07 OCT 2003) SET COST OFF FILE 'REGISTRY' ENTERED AT 15:18:00 ON 07 OCT 2003 E CHITOSAN/CN 1 S E3 L1E CHITOSAN 1498 S E3 L2L3 1497 S L2 NOT L1. 1433 S L3 NOT SQL/FA L4E TREHALOSE/CN 1 S E3 L5115 S 99-20-7/CRN L6 763 S 9012-76-4/CRN L7 18 0 S L6 AND L7 L91433 S L4,L7 FILE 'HCAPLUS' ENTERED AT 15:20:48 ON 07 OCT 2003 L10 12093 S L1 3160 S L9 L11 L1215211 S CHITOSAN L13 3172 S L3 15784 S L10-L13 L14L15 5999 S L5 167 S L6 L16 L17 8262 S TREHALOSE 8696 S L15-L17 L18 72 S L14 AND L18 L19 70 S L19 AND (PY<=1999 OR PRY<=1999 OR AYT<=1999) L20 E WORRALL E/AU L21 8 S E3, E4, E9 E ANHYDRO/PA,CS L22 35 S E3-E13 E WO2000-GB2254/AP, PRN E GB99-14412/AP, PRN L23 43 S L21, L22 0 S L23 AND L14 L24 3 S L23 AND L18 E PRESERVATION/CT E E3+ALL 2148 S E1 L26 11897 S E1+NT E E14+ALL 423 S E3 L28 601 S E3+NT E E2+ALL 5084 S E2 L30 42 S L18 AND L26 214 S L18 AND L27-L30 4 S L14 AND L26 L33 194 S L14 AND L27-L30 L34 8 S L31, L32 AND L33, L34 SEL DN AN 4 1 S L35 AND E1-E3 L36

4 S L25, L36

E FREEZE DRYING/CT

```
E E3+ALL
L38
           4716 S E12
                 E E15+ALL
          19277 S E2, E1+NT
L39
L40
            297 S L18 AND L38
L41
            139 S L18 AND L39
L42
             68 S L14 AND L38
L43
             20 S L14 AND L39
L44
              8 S .L19 AND L31-L34
L45
              5 S L19 AND L40-L43
L46
             12 S L44, L45
L47
              1 S L46 AND L37
L48
             11 S L46 NOT L47
                SEL DN AN 6 7
L49
              2 S E1-E6 AND L48
L50
              6 S L37, L47, L49
L51.
               6 S L50 AND L10-L50
                E DRYING/CT
                E E3+ALL
L52
          32988 S E2
                E E1+ALL
L53
            443 S E1
                E E6+ALL
L54
          19277 S E2, E1+NT
                E E13+ALL
          25489 S E6, E7, E5
L55
L56
            334 S L18 AND L52-L55
            133 S L14 AND L52-L55
L57
L58
              6 S L19 AND L56, L57
              3 S L51 AND L52-L57
L59
L60
              6 S L51, L59
L61
            637 S (BIOCHEM?(L)METHOD?)/SC, SX AND L14
L62
            649 S (BIOCHEM?(L)METHOD?)/SC, SX AND L18
             95 S L61, L62 AND L31-L34
L63
             22 S L63 AND L38-L41
L64
L65
             21 S L63 AND L52-L57
L66
             27 S L64, L65
L67
             16 S L66 AND (PY<=1999 OR PRY<=1999 OR AY<=1999)
             0 S L66 AND L19
L68
L69
             11 S L66 NOT L67
L70
             32 S L60, L67, L69
L71
             19 S L70 AND (PY<=1999 OR PRY<=1999 OR AY<=1999)
L72
             21 S L37, L71
L73
             11 S L70 NOT L72
                SEL HIT RN L70
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FILE 'REGISTRY' ENTERED AT 15:58:09 ON 07 OCT 2003 L74 2 S E1-E2

FILE 'REGISTRY' ENTERED AT 15:58:19 ON 07 OCT 2003

=> fil hcaplus FILE 'HCAPLUS' ENTERED AT 15:58:29 ON 07 OCT 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

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FILE COVERS 1907 - 7 Oct 2003 VOL 139 ISS 15 FILE LAST UPDATED: 6 Oct 2003 (20031006/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d 172 all hitstr tot

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L72 ANSWER 1 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN
```

AN . 2003:667367 HCAPLUS

DN 139:161828

TI Viable dried bacteria produced by drying in the presence of trehalose and divalent cation

IN Mateczun, Alfred J.; Peruski, Leonard F., Jr.

PA United States Dept. of the Navy, USA

SO U.S., 5 pp. CODEN: USXXAM

DT Patent

LA English

IC ICM A61K045-00

ICS A01N063-00; C12N001-00; C12N001-12; C12N001-20

NCL 435260000; 424093400; 424282100; 435252100; 435252330; 435822000; 435849000; 435879000; 435909000

CC 9-16 (Biochemical Methods)

Section cross-reference(s): 10

FAN.CNT 2

		_										
	PATENT NO.			DATE	•	APPLIC	CATION NO.	DATE				
				-								
ΡI	US	6610531	B1	20030826		US 199	98-159568	19980924	<			
	US	2003044965	A1	20030306		US 200	2-108344	20020329	<			
PRAI	US	1998-159568	B2	19980924	<							

AB A method is provided for preserving live bacteria by subjecting an aq. system contg. the growing bacteria to drying without special equipment, in the presence of trehalose with or without the addn. of divalent cations as stabilizing agents. Further, a dried compn. for preservation of aerobic bacteria in a viable state is provided. The dried compn. consists essentially of dried viable aerobic bacteria and an appropriate growth medium. The bacteria and growth medium are initially placed in an aq. soln. of 10 mM to 200 mM trehalose and a divalent cation, and dried at room temp.

ST viable dried bacteria drying trehalose divalent cation

IT Cations

(divalent; viable dried bacteria produced by drying in presence of trehalose and divalent cation)

IT Hydration, physiological

(rehydration; viable dried bacteria produced by drying in presence of trehalose and divalent cation)

IT Aerobic bacteria

Bacteria (Eubacteria)

Composition

Containers

Culture media

Drying

Escherichia coli

Gram-negative bacteria

Preservation

Preservation solutions (tissue)

Salmonella typhimurium Sealing Separation Shiqella flexneri Solutions Stabilizing agents Suspensions Temperature Vibrio cholerae Volume (viable dried bacteria produced by drying in presence of trehalose and divalent cation) 99-20-7, Trehalose 7646-85-7, Zinc chloride (ZnCl2), TΤ 7732-18-5, Water, biological studies 7786-30-3, biological studies Magnesium chloride (MgCl2), biological studies 10043-52-4, Calcium chloride (CaCl2), biological studies RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses) (viable dried bacteria produced by drying in presence of trehalose and divalent cation) THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD . RE.CNT RE (1) Anon; EP 0120111 A1 1984 (2) Goodrich; US 5800978 A 1998 HCAPLUS (3) Kosanke; US 5695541 A 1997 (4) Paau; US 4875921 A 1989 IT 99-20-7, Trehalose RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (viable dried bacteria produced by drying in presence of trehalose and divalent cation) RN 99-20-7 HCAPLUS .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME) CN

```
L72
    ANSWER 2 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN
AN
     2003:213042 HCAPLUS
DN
     138:242963
    Xerovac: an ultra rapid method for the dehydration and preservation of
ΤI
     live attenuated Rinderpest and Peste des Petits ruminants vaccines.
     [Erratum to document cited in CA135:231554]
ΑU
    Worrall, E. E.; Litamoi, J. K.; Seck, B. M.; Ayelet, G.
     Ty Mawr, Trefilan, Lampeter, Dyfed, SA48 8RD, UK
CS
SO
     Vaccine (2001), 19(28-29), 4086
     CODEN: VACCDE; ISSN: 0264-410X
PB
     Elsevier Science Ltd.
DT
     Journal
LA
     English
CC
     63-3 (Pharmaceuticals)
```

- AB On page 839, in Table 5, second column headed "Liq. virus pool titer + trehalose", the data on the first line should read 6.0 not 0.0.

 The cor. Table 5 is given.
- ST erratum dehydration preservation ruminant vaccine trehalose; dehydration, preservation ruminant vaccine trehalose erratum
- IT Bacteria (Eubacteria)
 Dehydration, physiological
 Freeze drying
 Peste-des-petits-ruminants virus
 Preservation

Rinderpest virus

Ruminant

Virus

(ultra rapid dehydration and preservation of live attenuated Rinderpest and Peste des Petits ruminants vaccines (Erratum))

IT 99-20-7, Trehalose

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (ultra rapid dehydration and preservation of live attenuated Rinderpest and Peste des Petits ruminants vaccines (Erratum))

IT 99-20-7, Trehalose

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(ultra rapid dehydration and preservation of live attenuated Rinderpest
and Peste des Petits ruminants vaccines (Erratum))

RN 99-20-7 HCAPLUS

CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

- L72 ANSWER 3 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN
- AN 2001:396599 HCAPLUS
- DN 135:2554
- TI Formulation of preservation mixtures containing sensitive biologicals to be stabilized for ambient temperature storage by drying
- IN Bronshtein, Victor; Linkowski, Lynn
- PA Universal Preservation Technologies, Inc., USA
- SO PCT Int. Appl., 34 pp. CODEN: PIXXD2
- DT Patent
- LA English
- IC ICM A01N001-02
- CC 9-11 (Biochemical Methods)
 Section cross-reference(s): 63

FAN.CNT 1

22111	PATENT NO.					ND	DATE			A	PPLI	CATI	и ис	0.	DATE	TE			
PI	WO 2001037656 WO 2001037656								W	20	00-U	61	20001122 <						
	,,,		AE,	AG,	AL,	AM,		AU,	AZ,										
			GD,	GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	ΚP,	KR,	KR,	

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KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ,
             NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SK, SL, TJ, TM, TR,
             TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU,
             TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                           20020821
                                           EP 2000-980766
                                                            20001122 <--
     EP 1231837
                       A2
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
                            20030415
                                           BR 2000-15738
                                                             20001122 <--
     BR 2000015738
                      · A
     JP 2003514556
                       Т2
                            20030422
                                           JP 2001-539285
                                                             20001122 <--
                                      <--
PRAI US 1999-166928P
                       Ρ
                            19991122
     WO 2000-US32261
                       W
                            20001122
     This invention relates to formulations and methods for preserving
AB
     sensitive biologicals, viruses, bacteria and eukaryotic cells by drying.
     More particularly, the invention relates to preservation mixts. comprising
     viruses or cells and protectants, including methylated monosaccharides,
     wherein the mixts. are adapted to stabilize these samples during
     dehydration and subsequent storage at ambient and higher temps.
     biol preservation preservative formulation drying storage
ST
     Whey
IT
        (albumins of; formulation of preservation mixts. contg. sensitive
        biologicals to be stabilized for ambient temp. storage by drying)
IT
     Health products
        (biologicals; formulation of preservation mixts. contg. sensitive
        biologicals to be stabilized for ambient temp. storage by drying)
ΙT
     Bovine respiratory syncytial virus
     Human parainfluenza virus 3
     Newcastle disease virus
     Streptococcus equi
        (drying and storage of; formulation of preservation mixts. contq.
        sensitive biologicals to be stabilized for ambient temp. storage by
        drying)
ΙT
     Bacteria (Eubacteria)
     Crystallization .
       Drying
     Eukaryote (Eukaryotae)
       Freeze drying
       Preservation
       Preservatives
     Prokaryote
     Storage
     Temperature effects, biological
     Virus
        (formulation of preservation mixts. contg. sensitive biologicals to be
        stabilized for ambient temp. storage by drying)
ΙT
     Albumins, biological studies
     Disaccharides
     Gelatins, biological studies
     Monosaccharides
     Polymers, biological studies
     Polyoxyalkylenes, biological studies
     Proteins, general, biological studies
     RL: BUU (Biological use, unclassified); NUU (Other use, unclassified);
     BIOL (Biological study); USES (Uses)
        (formulation of preservation mixts. contg. sensitive biologicals to be
        stabilized for ambient temp. storage by drying)
     Monosaccharides
IT
     RL: BUU (Biological use, unclassified); NUU (Other use, unclassified);
     BIOL (Biological study); USES (Uses)
        (methylated derivs.; formulation of preservation mixts. contg.
        sensitive biologicals to be stabilized for ambient temp. storage by
```

drying)

IT Oligosaccharides, biological studies

RL: BUU (Biological use, unclassified); NUU (Other use, unclassified);

BIOL (Biological study); USES (Uses)

(non-reducing derivs.; formulation of preservation mixts. contg.

sensitive biologicals to be stabilized for ambient temp. storage by drying)

IT Globulins, biological studies

RL: BUU (Biological use, unclassified); NUU (Other use, unclassified); BIOL (Biological study); USES (Uses)

(of whey; formulation of preservation mixts. contg. sensitive biologicals to be stabilized for ambient temp. storage by drying)

IT Drying

(spray; formulation of preservation mixts. contg. sensitive biologicals to be stabilized for ambient temp. storage by drying)

IT 9001-58-5, Isocitrate dehydrogenase 61969-99-1, luciferase

RL: PEP (Physical, engineering or chemical process); PROC (Process) (drying and storage of; formulation of preservation mixts. contg. sensitive biologicals to be stabilized for ambient temp. storage by drying)

IT 512-69-6, Raffinose

RL: BSU (Biological study, unclassified); BIOL (Biological study) (formulation of preservation mixts. contg. sensitive biologicals to be stabilized for ambient temp. storage by drying)

IT 57-50-1, Sucrose, biological studies 97-30-3, Methyl alpha.-glucopyranoside 99-20-7, Trehalose 142-47-2, Monosodium glutamate 709-50-2 9003-39-8, Pvp 12619-70-4, Cyclodextrin 25322-68-3, polyethylene glycol

stabilized for ambient temp. storage by drying)

RL: BUU (Biological use, unclassified); NUU (Other use, unclassified);

BIOL (Biological study); USES (Uses)
(formulation of preservation mixts. contq. sensitive biologicals to be

IT 99-20-7, Trehalose

RL: BUU (Biological use, unclassified); NUU (Other use, unclassified); BIOL (Biological study); USES (Uses)

(formulation of preservation mixts. contg. sensitive biologicals to be stabilized for ambient temp. storage by drying)

RN 99-20-7 HCAPLUS

CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

L72 ANSWER 4 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN

AN 2001:64128 HCAPLUS

DN 134:97541

TI Storage of microorganisms, cells and tissue

IN Codd, Anthony Arthur

PA Public Health Laboratory Service Board, UK

SO PCT Int. Appl., 33 pp. CODEN: PIXXD2

```
DT
     Patent
LA
     English
     ICM C12N005-00
TC
CC
     9-16 (Biochemical Methods)
     Section cross-reference(s): 10, 17
FAN.CNT 1
                      KIND DATE
                                           APPLICATION NO.
                                                            DATE
     PATENT NO.
                                           ______
     ______
                      ____
                           _____
                                                            _____
                                           WO 2000-GB2738 20000717 <--
     WO 2001005941
                     A2
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PI
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
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             HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
             LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
             SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
             YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
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             CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                           JP 2001-511155
                      Т2
                                                            20000717 <--
     JP 2003505024
                            20030212
                                           ZA 2002-340
                                                            20020115 <--
     ZA 2002000340
                       Α
                            20021010
PRAI GB 1999-16790
                       Α
                            19990716
                                     <--
     WO 2000-GB2738
                       W
                            20000717
AΒ
     A compn. for preserving viable microorganisms, cells or tissue comprises
     (a) a preservative combination of (i) a non-reducing disaccharide and (ii)
     a bulking agent; and (b) a buffer. Also described is a method of
     preserving viable microorganisms, cells or tissue, comprising combining
     viable microorganisms, cells or tissue with a preserving soln. comprising
     a non-reducing disaccharide, drying the combination to form a dried prepn.
     having a solids content of at least 80% by wt., and counting the viable
     microorganisms, cells or tissue in the dried prepn., whereby the dried
     prepn. can be combined with aq. buffer to yield an aq. prepn. comprising a
     predetd. count of viable microorganisms, cells or tissue.
ST
     storage microorganism cell tissue
ΙT
     Pressure
        (Atm.; storage of microorganisms, cells and tissue)
ΙT
     Proteins, general, biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (High mol. wt.; storage of microorganisms, cells and tissue)
ΙT
     Disaccharides
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (Non-reducing; storage of microorganisms, cells and tissue)
IT
     Carbohydrates, biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (Water-sol. polymeric; storage of microorganisms, cells and tissue)
ΙT
     Animal cell
        (mammalian; storage of microorganisms, cells and tissue)
IT
     Temperature
        (reduced; storage of microorganisms, cells and tissue)
IT
     Carbohydrates, biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (reducing sugars; storage of microorganisms, cells and tissue)
IT
     Albumins, biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (serum; storage of microorganisms, cells and tissue)
IT
     Air
     Animal tissue
     Bacteria (Eubacteria)
     Buffers
```

```
Cell
     Coloring materials
     Composition
      Dehydration
      Drying
     Drying agents
     Erythrocyte
     Escherichia coli
     Feces
     Food processing
     Fungi
     Microorganism
       Preservation
       Preservatives
     Protozoa
     Storage
     Urine
     Virus
        (storage of microorganisms, cells and tissue)
ΙT
    Albumins, biological studies
    Monosaccharides
     Noble gases, biological studies
     Silica gel, biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
    (Uses)
        (storage of microorganisms, cells and tissue)
IT
     Pharynx
        (swabs; storage of microorganisms, cells and tissue)
     7732-18-5, Water, biological studies 7782-44-7, Oxygen, biological
IT
     studies
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (storage of microorganisms, cells and tissue)
ΙT
     50-99-7, Glucose, biological studies
                                            57-50-1, Sucrose, biological
               63-42-3, Lactose 69-79-4, Maltose 99-20-7,
                111-30-8, Glutaraldehyde 528-50-7, Cellobiose
                                         9004-34-6D, Cellulose, Hydroxyalkyl,
     9004-32-4, Carboxymethylcellulose
     biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (storage of microorganisms, cells and tissue)
     99-20-7, Trehalose
IT
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (storage of microorganisms, cells and tissue)
     99-20-7 HCAPLUS
RN
     .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)
CN
```

```
2000:911307 HCAPLUS
ΑN
DN
     134:68451
     Enhanced stability and performance of cells and cell components
TΙ
     Potts, Malcolm; Helm, Richard
IN
    .Virginia Tech Intellectual Properties, Inc., USA
PA
SO
     PCT Int. Appl., 20 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
IC
     ICM C08B037-00
     ICS C07H001-08; C12P019-04; G01N001-30; G01N033-48
CC
     9-16 (Biochemical Methods)
FAN.CNT 1
                     KIND DATE
                                           APPLICATION NO.
     PATENT NO.
                                                            DATE
                      ----
                           -----
                                           -----
     ______
                            20001228
PΙ
     WO 2000078816
                     A1
                                           WO 2000-US16603 20000616 <--
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR,
             CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
             ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
             LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD,
             SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA,
             ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
             CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                                            20000616 <--
     EP 1203027
                           20020508
                                          EP 2000-942871
                       A1
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL
PRAI US 1999-139738P
                            19990618
                     Р
                                     <--
                       Ρ
     US 2000-178703P
                            20000128
                            20000616
     WO 2000-US16603
                      W
     The present invention provides a compn. of matter comprising a
AΒ
     substantially purified extracellular polysaccharide (EPS) from the
     terrestrial cyanobacterium Nostoc commune. When mixed with labile
     material (such as cells or cellular components) the EPS affords protection
     to the cells during desiccation and long-term storage. The invention thus
     provides an improved method for room temp. long-term storage of labile
     material.
     stability performance cell component
ST
ΙT
     Storage
        (Long-term; enhanced stability and performance of cells and cell
        components)
IT
     Carbohydrates, biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (Nonreducing; enhanced stability and performance of cells and cell
        components)
ΙT
     Heating
        (autoclaving; enhanced stability and performance of cells and cell
        components)
ΙT
     Agrochemicals
        (bio-; enhanced stability and performance of cells and cell components)
    Air
ΙT
    Animal cell
     Cell
     Composition
     Drugs
       Drying
     Eukaryote (Eukaryotae)
       Freeze drying
     Genetic engineering
     Mammal (Mammalia)
     Matter
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Mixing
    Nostoc commune
    Pesticides
      Preservation
    Prokaryote
    Purification
    Stability
    Vacuum
    Virus
        (enhanced stability and performance of cells and cell components)
    Alditols
IT
    Carbohydrates, biological studies
    Cyclitols
    Noble gases, biological studies
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (enhanced stability and performance of cells and cell components)
    Polysaccharides, biological studies
ΙT
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (exopolysaccharides; enhanced stability and performance of cells and
        cell components)
IT
    Animal tissue
        (mammalian; enhanced stability and performance of cells and cell
        components)
IT
    Alcohols, biological studies
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (polyhydric; enhanced stability and performance of cells and cell
        components)
IT
    Temperature
        (room; enhanced stability and performance of cells and cell components)
ΙT
    Drying
        (spray; enhanced stability and performance of cells and cell
        components)
IT
    Biological transport
        (uptake; enhanced stability and performance of cells and cell
        components)
     57-50-1, Sucrose, biological studies 99-20-7, Trehalose
ΙT
                                                 512-69-6, Raffinose
     470-55-3, Stachyose
                           470-57-5, Planteose
    597-12-6, Melezitose
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (enhanced stability and performance of cells and cell components)
    300823-68-1
ΙT
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (repeating unit, enhanced stability and performance of cells and cell
        components)
              THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
(1) Fischer; Planta Med 1993, V59(7), PA615
(2) Hill; J Appl Phycol, CAplus 1997:768211 1997, V9(3), P237 HCAPLUS
(3) Huang; J Phycol, CAplus 1999:69315 1998, V34(6), P962 HCAPLUS
IT
    99-20-7, Trehalose
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (enhanced stability and performance of cells and cell components)
     99-20-7 HCAPLUS
RN
     .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)
CN
```

L72 ANSWER 6 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN

AN 2000:874736 HCAPLUS

DN 135:231554

TI Xerovac: an ultra rapid method for the dehydration and preservation of live attenuated Rinderpest and Peste des Petits ruminants vaccines

AU Worrall, E. E.; Litamoi, J. K.; Seck, B. M.; Ayelet, G.

CS Ty Mawr, Trefilan, Lampeter, Dyfed, SA48 8RD, UK

SO Vaccine (2000), 19(7-8), 834-839 CODEN: VACCDE; ISSN: 0264-410X

PB Elsevier Science Ltd.

DT Journal

LA English

CC 63-3 (Pharmaceuticals)

AB The accepted procedure for the long-term preservation of live viruses and bacteria in vaccines has been lyophilization. We show that thermolabile viruses can be dehydrated in vitro, within 18 h, in an excipient contg. trehalose. We further demonstrate that in the resulting dehydrated state, where the viruses are captive in a metastable glass composed of trehalose, they are capable of resisting 45.degree.C for a period of 14 days with minimal loss of potency. The degree of thermotolerance achieved matches that of current 'thermostable' lyophilized vaccines, but with the distinct advantage of a shorter, cheaper and simpler process. The development and utilization of this process can make significant improvements in current live virus vaccine prodn. It presents a further step away from dependence on mandatory low temp. refrigerated storage and could lead to greater confidence in vaccine stability, potency and efficacy.

ST dehydration preservation ruminant vaccine *trehalose

IT Bacteria (Eubacteria)

Dehydration

Freeze drying

Peste-des-petits-ruminants virus

Preservation

Rinderpest virus

Ruminant

Virus

(ultra rapid dehydration and preservation of live attenuated Rinderpest and Peste des Petits ruminants vaccines)

IT 99-20-7, Trehalose

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (ultra rapid dehydration and preservation of live attenuated Rinderpest and Peste des Petits ruminants vaccines)

RE.CNT 12 THERE ARE 12 CITED REFERENCES AVAILABLE FOR THIS RECORD

- (1) Clegg, J; J Comp Biochem Physiol 1967, V20, P801 HCAPLUS
- (2) Colaco, C; Biotechnology 1992, Pl0
- (3) Coutinho, E; J Biotechnol 1988, V7, P23
- (4) Crowe, J; Cryobiology 1990, V27, P219 HCAPLUS
- (5) Franks, F; Biopharm 1991, V4, P38 HCAPLUS
- (6) Kinchin, I; Biologist 1995, V42, P4

```
(7) Levine, H; Biopharm 1992, V5, P36 HCAPLUS
```

- (8) Mariner, J; Vet Microbiol 1990, V21, P195 HCAPLUS
- (9) Reynolds, T; Adv Food Res 1965, V14, P167 MEDLINE
- (10) Rweyemamu, M; FAO Animal Production and Health Paper 1994, 118
- (11) Seki, K; Nature 1998, P395
- (12) Williams, R; Plant Physiol 1989, V89, P977

IT 99-20-7, Trehalose

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (ultra rapid dehydration and preservation of live attenuated Rinderpest and Peste des Petits ruminants vaccines)

RN 99-20-7 HCAPLUS

CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

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L72 ANSWER 7 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN
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AN 2000:790612 HCAPLUS

DN 133:319293

TI Method for the preservation of viruses and mycoplasma

IN Worrall, Eric Edward

PA UK

SO PCT Int. Appl., 24 pp. CODEN: PIXXD2

DT Patent

LA English

IC ICM C12N001-04

ICS C12N007-02; A61K039-155; A61K039-165; A61K039-20; A61K039-12; A61K039-13; A61K039-17

CC 9-11 (Biochemical Methods)

Section cross-reference(s): 10, 63

FAN.CNT 1

FAN.	:NT	1															•		
PATENT NO.				KI	ND	DATE			A	PPLI	CATI	ON NO	Ο.	DATE					
PI	PI WO 2000066710				A.	2	2000		W	O 20	00-G	B152	4	2000	0503	<			
_	WO	2000066710			A3 2001020			0208											
		W:	ΑE,	AL,	AM,	ÄΤ,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CR,	CU,	
			CZ,	DE,	DK,	DM,	ΕĒ,	ES,	FΙ,	GB,	GD,	GE,	GH,	GM,	HR,	ΗU,	ID,	IL,	
			IN,	IS,	JP,	KE,	KG,	ΚP,	KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	MA,	
			MD,	MG,	MK,	MN,	MW,	MX,	NO,	NZ,	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	
			SK,	SL,	ТJ,	TM,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VN,	YU,	ZA,	ZW,	AM,	
			AZ,	BY,	KG,	ΚZ,	MD,	RU,	ΤJ,	TM			·						
		RW:	GH,	GM,	KE,	LS,	MW,	SD,	SL,	SZ,	ΤZ,	UG,	ZW,	ΑT,	BE,	CH,	CY,	DE,	
			DK,	ES,	FI,	FR,	GB,	GR,	ΙE,	ΙT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,	
			CG,	CI,	CM,	GΑ,	GN,	GW,	ML,	MR,	NE,	SN,	TD,	ΤG					
	EP 1175486				A2 20020130					E	P 20	00-9	2743	8	20000503 <				
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,	
			ΙE,	SI,	LT,	LV,	FI,	RO											
	•			A					BR 2000-10249						0503	<			
JP 2002542815 PRAI GB 1999-9999			T	2	20021217			JP 2000-615735 20000503							<				
			A					<											

GB 1999-26698 A 19991112 <--WO 2000-GB1524 W 20000503

AB A biol.-active material comprising a live virus or mycoplasma is preserved by a method of desiccation, without lyophilization, in a matrix of glassy trehalose having a residual moisture content of not greater than 2%. The method comprises two vacuum drying stages. In a cycle time much shorter than a typical freeze drying process a virus or mycoplasma can be preserved to provide a material that can be rehydrated to give a vaccine having potency.

ST preservation virus mycoplasma

IT Drying

Freeze drying

Human poliovirus
Measles virus
Mixing
Mumps virus
Mycoplasma
Mycoplasma mycoides mycoides
Newcastle disease virus

Peste-des-petits-ruminants virus

Preservation

Pressure
Rinderpest virus
Rubella virus
Suspensions
Temperature
Vaccines
Vacuum
Virus
Yellow fever virus

(method for preservation of viruses and mycoplasma)

IT Hydration, physiological

(rehydration; method for preservation of viruses and mycoplasma)

IT 7732-18-5, Water, biological studies

RL: BSU (Biological study, unclassified); BIOL (Biological study) (method for preservation of viruses and mycoplasma)

IT 99-20-7, Trehalose

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(method for preservation of viruses and mycoplasma)

IT 99-20-7, Trehalose

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(method for preservation of viruses and mycoplasma)

RN 99-20-7 HCAPLUS

CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

```
ΑN
     2000:151414 HCAPLUS
DN
    132:205128
    A method for preserving mammalian organs removed
ΤI
IN
     Seki, Kunihiro
     Kanagawa University, Japan
PA
     Jpn. Kokai Tokkyo Koho, 6 pp.
SO
     CODEN: JKXXAF
DT
     Patent
LA
     Japanese
     ICM A01N001-02
IC
     9-11 (Biochemical Methods)
CC
FAN.CNT 1
                                          APPLICATION NO. DATE
     PATENT NO.
                     KIND DATE
                                          _____
     _____
                     ____
                                                           -----
                                          JP 1998-245052 19980831 <--
                      A2
                           20000307
     JP 2000072601
PΤ
PRAI JP 1998-245052
                           19980831 <--
    A method is provided for preserving mammalian organs removed (e.g., heart)
     for a long term. Mammalian organs removed are dehydrated using a
     dehydration agent (e.g., silica gel), soaked in a water- and oil-insol.
     inactive medium (e.g., perfluorocarbon, silicone oil), and maintained at a
     refrigeration temp. The effectiveness of this method was examinded with
     heart removed from rat by an electrophysiol. method i.e., ECG.
     organ preservation soln dehydration perfluorocarbon refrigeration
ST
ΙT
     Heart
        (ECG; method for preserving mammalian organs removed)
ΙT
    Dehydration
     Drying agents
     Heart
     Mammal (Mammalia)
     Organ, animal
       Organ preservation
       Preservation solutions (tissue)
     Rat
     Refrigeration
     Silica gel adsorbents
        (method for preserving mammalian organs removed)
IT
     Perfluorocarbons
     Polysiloxanes, biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (method for preserving mammalian organs removed)
     .99-20-7, Trehalose
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
    (Uses)
        (method for preserving mammalian organs removed)
IT
     99-20-7, Trehalose
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (method for preserving mammalian organs removed)
     99-20-7 HCAPLUS
RN
     .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)
CN
Absolute stereochemistry. Rotation (+).
```

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ANSWER 9 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN
L72
ΑN
     2000:98194 HCAPLUS
     132:133614
DN
     Agents containing sugar- or sugar alcohol-type surfactants and other
TI
     substances for preserving the freshness of cut flowers and vegetables
     Suzuki, Tadayuki; Kamei, Masatoshi; Hayashi, Masaharu; Kurita, Kazuhiko
ΙN
PA
     Kao Corporation, Japan
SO
     PCT Int. Appl., 45 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     Japanese
     ICM A01N003-02
IC
     5-3 (Agrochemical Bioregulators)
CC
     Section cross-reference(s): 17
FAN.CNT 1
                                                             DATE
     PATENT NO.
                      KIND
                            DATE
                                            APPLICATION NO.
                                            WO 1999-JP4080
                                                             19990729 <--
     WO 2000005946
                            20000210
                       A1
PT
         W: US
         RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
             PT, SE
                                            JP 1998-214106
                                                             19980729 <--
     JP 2000044401
                       A2
                            20000215
                                                             19981209 <--
     JP 2000169302
                       A2
                            20000620
                                            JP 1998-349965
     JP 2000103701
                       A2
                            20000411
                                            JP 1999-215861
                                                             19990729 <--
                                                            19990729 <--
     EP 1101402
                       `A1
                            20010523
                                            EP 1999-933160
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
PRAI JP 1998-214105
                       Α
                            19980729
                                       <--
     JP 1998-214106
                       .A
                            19980729
                                       <--
     JP 1998-349965
                       Α
                            19981209
                                      <--
     WO 1999-JP4080
                       W
                            19990729
                                      <--
     Highly safe agents for preserving the freshness of harvested plants such
AΒ
     as cut flowers and vegetables comprise a sugar- or sugar alc.-type
     surfactant together with .gtoreq.1 substance selected from among sugars,
     plant hormones, antioxidants, colloidal particle flocculating/pptg.
     agents, and microbicides and preservatives, preferably at a sp. wt. ratio.
     Thus, cut flowers (carnation, chrysanthemum, and rose) treated with an
     agent contg. 100 ppm sucrose fatty acid ester and 2.0% glucose lasted
     10-12 days, whereas flowers treated with 2.0% glucose alone lasted 5-6
     days and flowers in water lasted 3-5 days.
     preservative cut flower vegetable sugar surfactant
ST
IT
     Cut flower preservation
        (agents contg. sugar- or sugar alc.-type surfactants and other
        substances for)
IT
     Precipitation (chemical)
        (agents; preservatives for cut flowers and vegetables contg. sugar- or
```

sugar alc.-type surfactants and other substances)

RL: BAC (Biological activity or effector, except adverse); BSU (Biological

ΙT

Glycosides

IT

ΙT

IT

IT

TT

IT

TT

IT

IT

TT

study, unclassified); BUU (Biological use, unclassified); FFD (Food or feed use); BIOL (Biological study); USES (Uses) (alkyl polyglycosides; preservatives for cut flowers and vegetables contq. sugar- or sugar alc.-type surfactants and other substances) Hormones, plant RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BUU (Biological use, unclassified); FFD (Food or feed use); BIOL (Biological study); USES (Uses) (brassinosteroids; preservatives for cut flowers and vegetables contq. sugar- or sugar alc.-type surfactants and other substances) Food preservatives (contg. sugar- or sugar alc.-type surfactants and other substances) Preservatives (contg. sugar- or sugar alc.-type surfactants and other substances for keeping harvested plants fresh) Alditols Fatty acids, biological studies RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BUU (Biological use, unclassified); FFD (Food or feed use); BIOL (Biological study); USES (Uses) (esters; preservatives for cut flowers and vegetables contg. sugar- or sugar alc.-type surfactants and other substances) Amides, biological studies RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BUU (Biological use, unclassified); FFD (Food or feed use); BIOL (Biological study); USES (Uses) (fatty; preservatives for cut flowers and vegetables contq. sugar- or sugar alc.-type surfactants and other substances) Oligosaccharides, biological studies RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BUU (Biological use, unclassified); FFD (Food or feed use); BIOL (Biological study); USES (Uses) (fructose-lactose-contq.; preservatives for cut flowers and vegetables contg. sugar- or sugar alc.-type surfactants and other substances) Antimicrobial agents Antioxidants Carnation (Dianthus) Chinese cabbage Chrysanthemum Rose (Rosa) Spinach (Spinacia oleracea) Surfactants (preservatives for cut flowers and vegetables contg. sugar- or sugar alc.-type surfactants and other substances) Auxins Carbohydrates, biological studies Cytokinins Gibberellins Hormones, plant Polysaccharides, biological studies RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BUU (Biological use, unclassified); FFD (Food or feed use); BIOL (Biological study); USES (Uses) (preservatives for cut flowers and vegetables contg. sugar- or sugar alc.-type surfactants and other substances) Carbohydrates, biological studies RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BUU (Biological use, unclassified); FFD (Food or feed use); BIOL (Biological study); USES (Uses) (sugar esters; preservatives for cut flowers and vegetables contq. sugar- or sugar alc.-type surfactants and other substances) Amides, biological studies

ΙT

IT

TT

RN

CN

ΙT

RN

CN

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RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BUU (Biological use, unclassified); FFD (Food or
     feed use); BIOL (Biological study); USES (Uses)
        (sugar; preservatives for cut flowers and vegetables contg. sugar- or
        sugar alc.-type surfactants and other substances)
     9012-76-4, Chitosan
                         10043-01-3, Aluminum sulfate
     10043-52-4, Calcium chloride, uses
                                         147014-67-3, Kurifloc LC 541
     RL: NUU (Other use, unclassified); USES (Uses)
        (pptn. agent; preservatives for cut flowers and vegetables contq.
        sugar- or sugar alc.-type surfactants and other substances)
     50-70-4, Sorbitol, biological studies 50-99-7, D-Glucose, biological
             57-48-7, Fructose, biological studies 57-50-1, Sucrose,
                         57-50-1D, Sucrose, fatty acid esters 59-23-4,
     biological studies
     Galactose, biological studies 62-57-7, Aminoisobutyric acid
     Gibberellic acid 94-75-7, 2,4-D, biological studies 99-20-7,
                 148-24-3, 8-Hydroxyquinoline, biological studies
     Trehalose
                        1330-43-4, Sodium tetraborate
     525-79-1, Kinetin
                                                        1338-39-2, Rheodol
              7173-51-5, Didecyldimethylammonium chloride
                                                            13073-35-3,
                 23149-52-2, Silver thiosulfate
     Ethionine
                                                37266-93-6, DK Ester SL 18A
                  73904-70-8, Proxel 257285-60-2, Maidooru 10
     49669-74-1
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BUU (Biological use, unclassified); FFD (Food or
     feed use); BIOL (Biological study); USES (Uses)
        (preservatives for cut flowers and vegetables contg. sugar- or sugar
        alc.-type surfactants and other substances)
             THERE ARE 16 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
(1) Abbott Laboratories; US 5500403 A HCAPLUS
(2) Abbott Laboratories; AU 699897 B HCAPLUS
(3) Abbott Laboratories; EP 765114 A1 HCAPLUS
(4) Abbott Laboratories; WO 9534199 A1 HCAPLUS
(5) Abbott Laboratories; KR 97703697 A
(6) Abbott Laboratories; JP 10501553 A 1998
(7) Asahi Optical Co Ltd; JP 07187902 A 1995 HCAPLUS
(8) British Technology Group Ltd; ES 2113647 T3 HCAPLUS
(9) British Technology Group Ltd; AU 693092 B HCAPLUS
(10) British Technology Group Ltd; DE 69408664 A1
(11) British Technology Group Ltd; EP 696167 A1 HCAPLUS
(12) British Technology Group Ltd; WO 9424857 A1 HCAPLUS
(13) British Technology Group Ltd; JP 08509375 A 1996
(14) Kurorera Kogyo K K; JP 49120781 A 1974
(15) Riken Vitamin Oil Co Ltd; JP 5081856 A 1975
(16) T Hasegawa Co Ltd; JP 06336401 A 1994 HCAPLUS
     9012-76-4, Chitosan
     RL: NUU (Other use, unclassified); USES (Uses)
        (pptn. agent; preservatives for cut flowers and vegetables contq.
        sugar- or sugar alc.-type surfactants and other substances)
     9012-76-4 HCAPLUS
     Chitosan (8CI, 9CI)
                         (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     99-20-7, Trehalose
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BUU (Biological use, unclassified); FFD (Food or
     feed use); BIOL (Biological study); USES (Uses)
        (preservatives for cut flowers and vegetables contg. sugar- or sugar
        alc.-type surfactants and other substances)
     99-20-7 HCAPLUS
     .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)
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ANSWER 10 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN
ΑN
    2000:79265 HCAPLUS
    132:119565
DN
ΤI
    A method for manufacturing a dry enzymic analysis element with an improved
    preservative stability
    Makino, Yasuhiko; Muratani, Koji
IN
    Fuji Photo Film Co., Ltd., Japan
PΑ
    Jpn. Kokai Tokkyo Koho, 9 pp.
SO
    CODEN: JKXXAF
DT
    Patent
T.A
    Japanese
IC
    ICM G01N033-52
    ICS C12Q001-26; C12Q001-28; C12Q001-32; C12Q001-44; C12Q001-48;
         G01N033-92; G01N033-70
CC
    9-2 (Biochemical Methods)
FAN.CNT 1
                   KIND DATE
    PATENT NO.
                                          APPLICATION NO. DATE
     ______
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                                          ______
    JP 2000035427
                     A2
                           20000202
                                          JP 1998-203791 19980717 <--
PRAI JP 1998-203791
                           19980717 <--
    A method is described for manufg. a dry anal. element contg. an enzyme
    with an improved preservative stability for measuring a constituent in a
    body fluid (e.g., blood) or an other liq. sample. The dry anal. element
    comprises at least a color-developing reagent layer and a development
    layer formed on a transparent support material. An aq. soln. contg. at
    least enzyme(s) and disaccharide is supplied to the developing layer, and
    dried by blowing a warm air at about 30-70.degree.C. An improved
    preservative stability was obsd. by including a disaccharide selected from
    trehalose, sucrose, and maltose in the developing layer of the dry
    anal. element for measuring the total cholesterol using peroxidase,
    cholesterol esterase, and cholesterol oxidase. An improved preservative
    stability was also obsd. by including sucrose in the developing layer of
    the dry anal. element for measuring creatine kinase activity using
    hexokinase, glucose-6-phosphate dehydrogenase, and diaphorase.
ST ·
    dry enzymic analysis element preservation disaccharide
IT
    Analysis
        (enzymic anal.; method for manufg. dry enzymic anal. element with
        improved preservative stability)
IT
    Blood analysis
    Body fluid
      Drying
      Preservation
    Reflection spectroscopy
    Stabilizing agents
        (method for manufg. dry enzymic anal. element with improved
       preservative stability)
ΙT
    Disaccharides
```

RL: ARU (Analytical role, unclassified); ANST (Analytical study) (method for manufg. dry enzymic anal. element with improved

preservative stability)

IT Gelatins, uses

RL: DEV (Device component use); USES (Uses) (method for manufg. dry enzymic anal. element with improved preservative stability)

IT Polyesters, uses

RL: DEV (Device component use); USES (Uses) (method for manufg. dry enzymic anal. element with improved preservative stability)

9001-15-4, Kinase (phosphorylating), creatine
RL: ANT (Analyte); BAC (Biological activity or effector, except adverse);
BSU (Biological study, unclassified); ANST (Analytical study); BIOL
(Biological study)

(method for manufg. dry enzymic anal. element with improved preservative stability)

IT 67-07-2, Creatine phosphate 298-83-9 9001-40-5, Glucose-6-phosphate dehydrogenase 9001-51-8, Hexokinase 9003-99-0, Peroxidase 9026-00-0, Esterase, cholesterol 9028-76-6, Oxidase, cholesterol 13746-66-2, Potassium ferricyanide 37340-89-9, Diaphorase 94153-57-8
RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (method for manufg. dry enzymic anal. element with improved preservative stability)

IT 57-50-1, Sucrose, analysis 69-79-4, Maltose 99-20-7, Trehalose

Trenalose

RL: ARU (Analytical role, unclassified); ANST (Analytical study) (method for manufg. dry enzymic anal. element with improved preservative stability)

IT 25038-59-9, Polyethylene terephthalate, uses RL: DEV (Device component use); USES (Uses)

(method for manufg. dry enzymic anal. element with improved preservative stability)

IT 13463-67-7, Titanium oxide, uses

RL: NUU (Other use, unclassified); USES (Uses)

(method for manufg. dry enzymic anal. element with improved preservative stability)

IT 57-88-5, Cholesterol, analysis

RL: ANT (Analyte); ANST (Analytical study)
(total; method for manufg. dry enzymic anal. element with improved preservative stability)

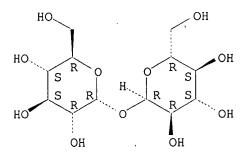
IT 99-20-7, Trehalose

RL: ARU (Analytical role, unclassified); ANST (Analytical study) (method for manufg. dry enzymic anal. element with improved preservative stability)

RN 99-20-7 HCAPLUS

CN ..alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



L72 ANSWER 11 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN AN 2000:35902 HCAPLUS

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DN
     132:339192
     Development of a freeze-dried formulation of insulin-loaded
     chitosan nanoparticles intended for nasal administration
     Fernandez-Urrusuno, R.; Romani, D.; Calvo, P.; Vila-Jato, J. L.; Alonso,
ΑU
     Department of Pharmaceutical Technology, School of Pharmacy, Santiago de
CS
     Compostela, 15706, Spain
     S.T.P. Pharma Sciences (1999), 9(5), 429-436
SO
     CODEN: STSSE5; ISSN: 1157-1489
PΒ
     Editions de Sante
DT
     Journal
LA
     English
CC
     63-6 (Pharmaceuticals)
AΒ
     The aim of this work was to develop a freeze-dried formulation of
     insulin-loaded chitosan nanoparticles and to assess their
     efficacy for reducing the plasma glucose levels after masal
     administration. Chitosan nanoparticles were prepd. by
     ionotropic gelation of chitosan with tripolyphosphate anions and
     then characterized in vitro (size, zeta potential, insulin loading and
     release) and in vivo (plasma glucose levels). The nanoparticles were also
     freeze-dried in the presence of various cryoprotective agents and then
     characterized upon reconstitution in water. Fresh chitosan
     nanoparticles displayed a pos. charge and a high insulin loading (up to
     55%). In vitro release studies showed that insulin was totally released
     from the nanoparticles in <1 h. Upon freeze-drying in the presence of
     trehalose or sucrose, the nanoparticles were freely reconstituted
     in water without a significant change in their size, zeta potential,
     insulin loading and release rate. The in vivo evaluation in conscious
     rabbits revealed that insulin-assocd. nanoparticles are able to reduce the
     plasma glucose levels to a greater extent than insulin chitosan
     solns. Furthermore, the in vivo efficacy of the reconstituted
     insulin-loaded nanoparticles administered intranasally was fully
     maintained. Consequently, freeze-dried chitosan nanoparticles
     can be proposed as useful vehicles for increasing the nasal absorption of
     insulin.
ST
     insulin chitosan nanoparticle nasal freeze dried
     Dissolution rate
     Drug bioavailability
       Freeze drying
     Gelation
     Particle size distribution
     Zeta potential
        (freeze-dried formulation of insulin-loaded chitosan
        nanoparticles for nasal administration)
ΙT
     Drug delivery systems
        (nanoparticles; freeze-dried formulation of insulin-loaded
        chitosan nanoparticles for nasal administration)
     Drug delivery systems
IT
        (nasal; freeze-dried formulation of insulin-loaded chitosan
        nanoparticles for nasal administration)
     9004-10-8, Insulin, biological studies
ΙT
     RL: BAC (Biological activity or effector, except adverse); BPR (Biological
     process); BSU (Biological study, unclassified); THU (Therapeutic use);
     BIOL (Biological study); PROC (Process); USES (Uses)
       (freeze-dried formulation of insulin-loaded chitosan
        nanoparticles for nasal administration)
     57-50-1, Sucrose, processes 99-20-7, Trehalose
     RL: PEP (Physical, engineering or chemical process); PROC (Process)
       (freeze-dried formulation of insulin-loaded chitosan
        nanoparticles for nasal administration)
                          14127-68-5, Tripolyphosphate
IT
     9012-76-4, Chitosan
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (freeze-dried formulation of insulin-loaded chitosan
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nanoparticles for nasal administration)

RE.CNT 22 THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

- (1) Almeida, A; J Pharm Pharmacol 1993, V45, P198 HCAPLUS
- (2) Alpar, H; J Drug Targeting 1994, V2, P147 MEDLINE
- (3) Artursson, P; Pharm Res 1994, V11, P1358 HCAPLUS
- (4) Aspden, T; Eur J Pharm Sci 1996, V4, P23 HCAPLUS
- (5) Calvo, P; J Appl Pol Sci 1997, V63, P125 HCAPLUS
- (6) Calvo, P; Pharm Res 1997, V14, P1431 HCAPLUS
- (7) Calvo, P; Proceed 2nd World Meeting APGI/APV 1998, P1111
- (8) Dondeti, P; Pharmazie 1994, V49, P505 HCAPLUS
- (9) Fernandez-Urrusuno, R; Pharm Res 1999, V16, P1576 HCAPLUS
- (10) Her, L; Pharm Res 1994, V11, P54 HCAPLUS
- (11) Illum, L; Pharm Res 1994, V11, P1186 HCAPLUS
- (12) Illum, L; Pharm Res 1998, V15, P1326 HCAPLUS
- (13) Jin, C; Trans Tianjin Univ 1996, V2, P25
- (14) Lehr, C; Int J Pharm 1992, V78, P43 HCAPLUS
- (15) Li, Y; STP Pharma Sci 1991, V1, P363 HCAPLUS
- (16) Lin, S; Chem Pharm Bull 1992, V40, P2491 HCAPLUS
- (17) Luessen, H; Pharm Res 1996, V13, P1668 HCAPLUS
- (18) Mayor, S; Int J Pharm 1997, V149, P123 HCAPLUS
- (19) Ohya, Y; J M S Pure Appl Chem 1994, VA31, P629 HCAPLUS
- (20) Remunan-Lopez, C; Eur J Pharm Biopharm 1998, V45, P49 HCAPLUS
- (21) Tobio, M; Pharm Res 1998, V15, P274
- (22) Uchegbu, I; J Pharm Pharmacol 1998, V50, P453 HCAPLUS
- IT 99-20-7, Trehalose

RL: PEP (Physical, engineering or chemical process); PROC (Process) (freeze-dried formulation of insulin-loaded chitosan nanoparticles for nasal administration)

RN 99-20-7 HCAPLUS

CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

IT 9012-76-4, Chitosan

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (freeze-dried formulation of insulin-loaded chitosan nanoparticles for nasal administration)

RN 9012-76-4 HCAPLUS

CN Chitosan (8CI, 9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

- L72 ANSWER 12 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN
- AN 1999:778625 HCAPLUS
- DN 132:177426
- TI Combined effects of **trehalose** and cations on the thermal resistance of .beta.-galactosidase in freeze-dried systems
- AU Mazzobre, M. F.; Del Pilar Buera, M.
- CS Facultad de Ciencias Exactas y Naturales, Departamento de Industrias, Ciudad Universitaria, Universidad de Buenos Aires, Buenos Aires, 1428,

SO Biochimica et Biophysica Acta (1999), 1473(2-3), 337-344 CODEN: BBACAQ; ISSN: 0006-3002 PB Elsevier Science B.V. DT Journal LAEnglish CC 7-8 (Enzymes) Section cross-reference(s): 9 The purpose of this study was to investigate the combined effects of AB trehalose and cations on the preservation of .beta.-galactosidase in freeze-dried systems and their relationship to phys. properties. Differential scanning calorimetry was employed to measure the glass transition temp. (Tg) and the endothermal peak area, related to the amt. of cryst. trehalose dihydrate present in the samples. In systems in which the trehalose matrix was humidified to conditions which allowed a high proportion of trehalose to crystallize, the enzyme was rapidly inactivated upon heating at 70.degree.. In these conditions the addn. of CsCl, NaCl and particularly KCl or MgCl2, improved the enzyme stability with respect to that obsd. in matrixes contg. only trehalose. For a given moisture content, addn. of salts produced very little change on the glass transition temp.; therefore the protective effect could not be attributed to a higher Tg value. The crystn. of trehalose dihydrate in the humidified samples was delayed in the trehalose/salt systems (principally in the presence of Mg2+) and a parallel improvement of enzyme stability was obsd. trehalose salt freeze drying enzyme stability; beta galactosidase preservation freeze drying trehalose; thermal stability protein freeze drying trehalose Freeze drying Hydration, chemical Preservation Thermal stability (combined effects of trehalose and cations on thermal resistance of .beta.-galactosidase in freeze-dried systems) IT Enzymes, properties Proteins, general, properties RL: PEP (Physical, engineering or chemical process); PRP (Properties); PROC (Process) (combined effects of trehalose and cations on thermal resistance of .beta.-galactosidase in freeze-dried systems) 7447-40-7, Potassium chloride IT 99-20-7, Trehalose 7647-14-5, Sodium chloride, properties 7647-17-8, (KCl), properties Cesium chloride (CsCl), properties 7786-30-3, Magnesium chloride (MgCl2), properties RL: NUU (Other use, unclassified); PRP (Properties); USES (Uses) (combined effects of trehalose and cations on thermal resistance of .beta.-galactosidase in freeze-dried systems) IT 9031-11-2, .beta.-Galactosidase RL: PEP (Physical, engineering or chemical process); PRP (Properties); PROC (Process) (combined effects of trehalose and cations on thermal resistance of .beta.-galactosidase in freeze-dried systems) THERE ARE 38 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT 38 (1) Aldous, B; Cryo-Lett 1995, V16, P181 HCAPLUS (2) Angyal, S; Aust J Chem 1972, V25, P1957 HCAPLUS (3) Anon; Handbook of Chemistry and Physics, 67th edn 1986-1987 (4) Anon; Merck Index, 12th edn (5) Buera, M; Food Res Int 1995, V28, P359 HCAPLUS (6) Cardona, S; J Food Sci 1997, V62, P105 HCAPLUS (7) Carpenter, J; Arch Biochem Biophys 1986, V250, P505 HCAPLUS

(8) Carpenter, J; Biochim Biophys Acta 1987, V923, P109 HCAPLUS

- (9) Carpenter, J; Cryobiology 1988, V25, P459 HCAPLUS
- (10) Chan, H; Pharm Res 1996, V13, P756 HCAPLUS
- (11) Colaco, C; Bio/Technology 1992, V10, P1007 HCAPLUS
- (12) Cook, W; Carbohydr Res 1973, V31, P265 HCAPLUS
- (13) Crowe, J; Annu Rev Physiol 1998, V60, P73 HCAPLUS
- (14) Crowe, J; Cryobiology 1994, V31, P355 MEDLINE
- (15) Crowe, L; Biophys J 1996, V71, P2087 HCAPLUS
- (16) Crowe, L; Science 1984, V233, P701
- (17) Dickson, R; J Bacteriol 1979, V137, P51 HCAPLUS
- (18) Green, J; J Phys Chem 1989, V93, P2880 HCAPLUS
- (19) Guy, E; J Dairy Sci 1978, V61, P147 HCAPLUS
- (20) Leslie, S; Appl Environ Microbiol 1995, V61, P3592 HCAPLUS
- (21) Mazzobre, M; Lebensm-Wiss-Technol 1997, V30, P324 HCAPLUS
- (22) Miller, D; Pharm Res 1997, V14, P578 HCAPLUS
- (23) Miller, D; Pharm Res 1998, V15, P1215 HCAPLUS
- (24) Morel-Desrosier, N; J Chem Soc Faraday Trans 1991, V87, P2173
- (25) Palumbo, M; J Food Sci 1995, V60, P117
- (26) Rongere, P; J Chem Soc Faraday Trans 1995, V91, P2771 HCAPLUS
- (27) Roos, Y; Biotechnol Prog 1991, V7, P49 HCAPLUS
- (28) Roos, Y; Carbohydr Res 1993, V238, P39 HCAPLUS
- (29) Roos, Y; J Food Sci 1991, V56, P1676 HCAPLUS
- (30) Schebor, C; Biotechnol Prog 1997, V13, P857 HCAPLUS
- (31) Shafizadeh, F; J Org Chem 1973, V28, P3710
- (32) Shamblin, S; J Therm Anal 1996, V47, P1567 HCAPLUS
- (33) Sussich, F; J Am Chem Soc 1998, V120, P7893 HCAPLUS
- (34) Suzuki, T; J Chem Eng Jpn 1997, V30, P609 HCAPLUS
- (35) Tan, C; Cryobiology 1995, V32, P60
- (36) Taylor, L; J Pharm Sci 1998, V87, P347 HCAPLUS
- (37) Voget, C; Lebensm-Wiss-Technol 1994, V27, P324 HCAPLUS
- (38) Whitaker, J; Food Preservation by Moisture Control 1995, P255 HCAPLUS
- IT 99-20-7, Trehalose
 - RL: NUU (Other use, unclassified); PRP (Properties); USES (Uses) (combined effects of trehalose and cations on thermal

resistance of .beta.-galactosidase in freeze-dried systems)

- RN 99-20-7 HCAPLUS
- CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

- L72 ANSWER 13 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN
- AN 1999:702928 HCAPLUS
- DN 132:32848
- TI Preservation of frozen yeast cells by trehalose
- AU Diniz-Mendes, L.; Bernardes, E.; de Araujo, P. S.; Panek, A. D.; Paschoalin, V. M. F.
- CS Departamento de Bioquimica, Instituto de Quimica, Universidade Federal do Rio de Janeiro, Rio de Janeiro, 21949-900, Brazil
- SO Biotechnology and Bioengineering (1999), 65(5), 572-578 CODEN: BIBIAU; ISSN: 0006-3592
- PB John Wiley & Sons, Inc.

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DT
     Journal
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English LA

CC 9-11 (Biochemical Methods)

Two different methods commonly used to preserve intact yeast AB cells-freezing and freeze-drying-were compared. Different yeast cells . submitted to these treatments were stored for 28 days and cell viability assessed during this period. Intact yeast cells showed to be less tolerant to freeze-drying than to freezing. The rate of survival for both treatments could be enhanced by exogenous trehalose (10%) added during freezing and freeze-drying treatments or by a combination of two procedures: a pre-exposure of cells to 40.degree.C for 60 min and addn. of trehalose. A max. survival level of 71.5 .+-. 6.3% after freezing could be achieved at the end of a storage period of 28 days, whereas only 25.0 .+-. 1.4% showed the ability to tolerate freeze-drying treatment, if both low-temp. treatments were preceded by a heat exposure and addn. of trehalose to yeast cells. Increased survival ability was also obtained when the pre-exposure treatment of yeast cells was performed at 10.degree.C for 3 h and trehalose was added: these treatments enhanced cell survival following freezing from 20.5 .+-. 7.7% to 60.0 .+-. 3.5%. Although both mild cold and heat shock treatments could enhance cell tolerance to low temp., only the heat treatment was able to increase the accumulation of intracellular trehalose whereas, during cold shock exposure, the intracellular amt. of trehalose remained unaltered. Intracellular trehalose levels seemed not to be the only factor contributing to cell tolerance against freezing and freeze-drying treatments; however, the protection that this sugar confers to cells can be exerted only if it is to be found on both sides of the plasma membrane.

cryopreservation yeast trehalose cryoprotectant; preservation ST yeast freezing freeze drying trehalose cryoprotectant

IT Temperature effects, biological

(cold, effect of mild cold shock; preservation of frozen yeast cells by trehalose)

Temperature effects, biological IT

(heat, effect of mild heat shock; preservation of frozen yeast cells by trehalose)

IT Cryopreservation

Cryoprotectants

Freeze drying

Freezing

Saccharomyces cerevisiae

(preservation of frozen yeast cells by trehalose)

IT Biological transport

(uptake, of trehalose; preservation of frozen yeast cells by trehalose)

ΙT 99-20-7, Trehalose

RL: BPR (Biological process); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); PROC (Process); USES (Uses)

(preservation of frozen yeast cells by trehalose).

THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT RE

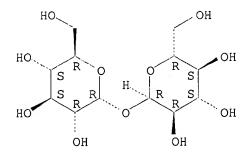
- (1) Bell, W; Eur J Biochem 1992, V209, P951 HCAPLUS
- (2) Carpenter, J; Biochim Biophys Acta 1987, V923, P109 HCAPLUS
- (3) Coutinho, C; J Biotechnol 1988, V7, P23 HCAPLUS
- (4) Crowe, J; Cryobiology 1990, V27, P219 HCAPLUS
- (5) Crowe, L; Biochim Biophys Acta 1986, V861, P131 HCAPLUS
- (6) Hottiger, T; FEBS Lett 1987, V220, P113 HCAPLUS (7) Kaul, S; Cell Mol Biol 1992, V38, P553 HCAPLUS
- (8) Komatsu, Y; Ferment Res Inst 1987, V68, P27
- (9) Leslie, S; Appl Environ Microbiol 1995, V61, P3592 HCAPLUS
- (10) Lindquist, S; Annu Rev Genet 1988, V22, P631 HCAPLUS
- (11) Obuchi, K; Cryo-Lett 1990, V11, P287

- (12) Pellerin-Mendes, C; Cryobiology 1997, V35, P173 HCAPLUS
- (13) Prestrelski, S; Arch Biochem Biophys 1993, V303, P465 HCAPLUS
- (14) Sanchez, Y; EMBO J 1992, V11, P2357 HCAPLUS
- (15) Stambuk, B; Biochim Biophys Acta 1998, V1379, P118 HCAPLUS
- (16) Sussman, A; Nature 1959, V130, P1343 HCAPLUS
- (17) Tatchell, K; Nature 1984, V309, P523 HCAPLUS
- (18) Tourinho dos Santos, C; Braz J Med Biol Res 1994, V27, P627 HCAPLUS
- (19) Van Laere, A; FEMS Microbiol Rev 1989, V63, P201 HCAPLUS
- (20) Zahringer, H; FEBS Lett 1997, V412, P615 MEDLINE
- IT 99-20-7, Trehalose

RL: BPR (Biological process); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); PROC (Process); USES (Uses)

(preservation of frozen yeast cells by trehalose)

- RN 99-20-7 HCAPLUS
- CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)



- L72 ANSWER 14 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN
- AN 1999:581741 HCAPLUS
- DN 132:148676
- TI Freeze-dried formulation for direct 99mTc-labeling ior-egf/r3 MAb. Additives, biodistribution, and stability
- AU Morales, A. A. M.; Nunez-Gandolff, G.; Perez, N. P.; Veliz, B. C.; Caballero-Torres, I.; Duconge, J.; Fernandez, E.; Crespo, F. Z.; Veloso, A.; Iznaga-Escobar, N.
- CS Center of Molecular Immunology, Havana, Cuba
- SO Nuclear Medicine and Biology (1999), 26(6), 717-723 CODEN: NMBIEO; ISSN: 0969-8051
- PB Elsevier Science Inc.
- DT Journal
- LA English
- CC 9-10 (Biochemical Methods)
- Monoclonal antibodies (MAbs) were useful for immunoscintigraphic AB applications in clin. diagnosis since they were introduced in nuclear medicine practice. The MAb ior egf/r3 developed at the Center of Mol. Immunol. (Havana, Cuba) is a murine antibody that recognizes the human epidermal growth factor receptor (EGF-R) and was used widely in the radioimmunodiagnosis of tumors of epithelial origin. Based on the direct Schwarz method, the present report describes the prepn. of a freeze-dried formulation for radiolabeling the MAb ior egf/r3 with 99mTc for immunoscintigraphic applications. Radiolabeling efficiency, effects on immunoreactivity, biodistribution, pharmacokinetic, and stability of the formulation are reported. The study demonstrated that the freeze-dried formulation can be labeled with 99mTc at high yield. The resulting 99mTc-labeled ior egf/r3 MAb can be used to visualize in vivo human tumors of epithelial origin by immunoscintigraphy studies. The kit does not need any other addn. or purifn. at the time of tagging other than the requisite amt. of pertechnetate (40-50 mCi). Because the contents of the kit are

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lyophilized, no special storage or transportation is required.
ST
     freeze dried technetium 99m labeled ioregfr3 monoclonal antibody
     immunoscintigraphy; EGFR lyophilization technetium 99m labeled ioregfr3
     monoclonal antibody immunoscintigraphy; epidermal growth factor receptor
     ioregfr3 antibody lyophilization immunoscintigraphy
ΙT
     Freeze drying
        (EGF-R detected by lyophilized 99mTc-labeling ior-eqf/r3 MAb using
        immunoscintigraphy)
ΙT
     Epidermal growth factor receptors
     RL: ANT (Analyte); THU (Therapeutic use); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
        (EGF-R detected by lyophilized 99mTc-labeling ior-egf/r3 MAb using
        immunoscintigraphy)
ΙT
     Scintigraphy
        (immunoscintigraphy; lyophilized 99mTc-labeling ior-egf/r3 MAb for
        immunoscintigraphy)
ΙT
     Brain
     Intestine
     Kidney
     Liver
     Lung
     Pancreas
     Spleen
     Stomach
        (lyophilized 99mTc-labeling ior-egf/r3 MAb for immunoscintigraphy,
        biodistribution)
ΙT
     Antibodies
     RL: BPN (Biosynthetic preparation); BPR (Biological process); BSU
     (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological
     study); PREP (Preparation); PROC (Process); USES (Uses)
        (monoclonal; lyophilized 99mTc-labeling ior-egf/r3 MAb for
        immunoscintigraphy)
     Physiological saline solutions
IT
        (phosphate-buffered; additives to lyophilized 99mTc-labeling ior-egf/r3
        MAb for immunoscintigraphy)
IT
     Colloids
       Preservatives
        (preservatives effect on stability of lyophilized 99mTc-labeling
        ior-egf/r3 MAb for immunoscintigraphy)
     50-70-4, D-Glucitol, analysis
                                    50-99-7, Glucose, analysis
ΙT
                                                                   57-50-1,
                69-65-8, Mannitol
                                   87-89-8, Inositol 99-20-7,
     analysis
     Trehalose
     RL: ARU (Analytical role, unclassified); THU (Therapeutic use); ANST
    (Analytical study); BIOL (Biological study); USES (Uses)
        (additives to lyophilized 99mTc-labeling ior-egf/r3 MAb for
        immunoscintigraphy)
     14133-76-7, Technetium 99, analysis
ΙT
     RL: ARU (Analytical role, unclassified); THU (Therapeutic use); ANST
     (Analytical study); BIOL (Biological study); USES (Uses).
        (metastable; EGF-R detected by lyophilized 99mTc-labeling ior-egf/r3
        MAb using immunoscintigraphy)
ΙT
     67-64-1, Acetone, analysis
     RL: ARU (Analytical role, unclassified); THU (Therapeutic use); ANST
     (Analytical study); BIOL (Biological study); USES (Uses)
        (preservatives effect on stability of lyophilized 99mTc-labeling
        ior-eqf/r3 MAb for immunoscintigraphy)
              THERE ARE 39 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
(1) Bale, W; Cancer Res 1980, V40, P2965 HCAPLUS
(2) Carpenter, J; Cryobiology 1988, V25, P459 HCAPLUS
(3) Cohen, S; J Biol Chem 1982, V257, P1523 HCAPLUS
(4) Cohen, S; Proc Natl Acad Sci USA 1982, V79, P6237 HCAPLUS
(5) Colcher, D; Cancer Res 1987, V47, P1185 MEDLINE
```

- (6) Crowe, J; Biochem J 1987, V242, P2
- (7) Crowe, J; Cryobiology 1990, V27, P219 HCAPLUS
- (8) Divgi, C; J Natl Cancer Inst 1991, V83, P97 MEDLINE
- (9) Draber, P; J Immunol Methods 1995, V181, P37 HCAPLUS
- (10) Ellman, G; Arch Bioch Biophys 1959, V82, P70 HCAPLUS(11) Fernandez, A; Interferon y Biotecnologia 1989, V6, P289
- (12) Fernandez, A; Interferon y Biotechologia 1989, V6, F288 (12) Fernandez, A; J Cell Biochem 1992, V49, P157 HCAPLUS
- (13) Goldenberg, D; Cancer Res 1990, V50, P909
- (14) Goldenberg, D; Semin Nucl Med 1989, V19, P262 MEDLINE
- (15) Greenwood, F; Biochem J 1963, V89, P114 MEDLINE
- (16) Israeli, E; Cryobiology 1993, V30, P519 HCAPLUS
- (17) Iznaga-Escobar, N; Nucl Med Biol 1996, V23, P635 MEDLINE
- (18) Iznaga-Escobar, N; Nucl Med Biol 1996, V23, P641 MEDLINE
- (19) Jennings, T; IVD Technology 1997, VJanuary/February, P38
- (20) Lindmo, T; J Immunol Methods 1984, V72, P77 HCAPLUS
- (21) Macias, A; Interferon y Biotecnologia 1985, V2, P115
- (22) Mather, S; J Nucl Med 1990, V31, P692 HCAPLUS
- (23) Morales, A; Nucl Med Biol 1998, V25, P25 HCAPLUS
- (24) Morales, A; Nucl Med Biol 1999, V26, P275
- (25) Morrison, R; Int J'Nucl Med Biol 1984, V11, P184 MEDLINE
- (26) Murray, J; J Nucl Med 1987, V28, P25 MEDLINE
- (27) Neal, D; Lancet 1985, V1, P346
- (28) Perez, R; Breast Cancer Res Treat 1984, V4, P189 HCAPLUS
- (29) Pikal, M; Biopharm 1990, V3, P18 HCAPLUS
- (30) Reilly, R; Nucl Med Commun 1993, V14, P347 MEDLINE
- (31) Rios, M; Anticancer Res 1992, V12, P205 MEDLINE
- (32) Rodriguez, N; Rev Esp Med Nucl. 1995, V14, P285
- (33) Roser, B; Biopharm 1991, V5, P47
- (34) Rudolph, A; Cryobiology 1990, V27, P585 HCAPLUS
- (35) Schwarz, A; J Nucl Med 1987, V28, P721
- (36) Singh, A; Nucl Med Biol 1994, V21, P277 HCAPLUS.
- (37) Snelling, L; Hybridoma 1995, V14, P111 MEDLINE
- (38) Thrall, J; J Nucl Med 1976, V19, P796
- (39) Veale, D; Br J Cancer 1987, V55, P513 MEDLINE
- IT 99-20-7, Trehalose

RL: ARU (Analytical role, unclassified); THU (Therapeutic use); ANST
(Analytical study); BIOL (Biological study); USES (Uses)
 (additives to lyophilized 99mTc-labeling ior-egf/r3 MAb for
 immunoscintigraphy)

RN 99-20-7 HCAPLUS

CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

- L72 ANSWER 15 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN
- AN 1999:462706 HCAPLUS
- DN 131:115668
- TI Keeping freshness of flowers and vegetables using trehalose and chitosan
- IN Inoue, Tadashi; Inoue, Takeshi

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PA Okayama Oyo Kagaku Y. K., Japan
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SO Jpn. Kokai Tokkyo Koho, 7 pp.

CODEN: JKXXAF

DT Patent

LA Japanese

IC ICM A23B007-153

ICS A01G007-00; A01N063-00; C08L005-08

CC 17-4 (Food and Feed Chemistry)

FAN.CNT 1

PATENT NO. KIND DATE APPLICATION NO. DATE

PI JP 11196765 A2 19990727 JP 1998-8604 19980120 <--

PRAI JP 1998-8604 19980120 <--

AB Freshness of plants, e.g. flowers, vegetables, is kept by soaking, coating, or spraying the plants with solns. contg. 1-5% trehalose, enzymically prepd. from maltose, and 0.1-0.0001% water-sol. chitosan showing pH 6.5-7.6. Trehalose and

chitosan control rate of flowering and formation of floral axis. Cut roses were soaked in an aq. soln. (pH 7) contg. 5% trehalose and 0.01% water-sol. low-mol.-wt. chitosan to be significantly prevented from wilting and to flower slowly.

ST plant freshness preservation trehalose chitosan soln; flower vegetable freshness preservation trehalose chitosan

IT Cut flower preservation

Flower

Food preservation

Food preservatives

Plant (Embryophyta)

Vegetable

(keeping freshness of flowers and vegetables using solns. of trehalose and chitosan)

IT 99-20-7, Trehalose 9012-76-4, Chitosan

RL: AGR (Agricultural use); FFD (Food or feed use); BIOL (Biological study); USES (Uses)

(keeping freshness of flowers and vegetables using solns. of trehalose and chitosan)

IT 99-20-7, Trehalose 9012-76-4, Chitosan

RL: AGR (Agricultural use); FFD (Food or feed use); BIOL (Biological study); USES (Uses)

(keeping freshness of flowers and vegetables using solns. of trehalose and chitosan)

RN 99-20-7 HCAPLUS

CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 9012-76-4 HCAPLUS

CN Chitosan (8CI, 9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

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L72 ANSWER 16 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN
     1999:286756 HCAPLUS
ΑN
     131:70761
DN
     A stabilized reaction mixture for in vitro translation of mRNA
TI
     Shaloiko, L. A.; Gorokhovhtskii, A. Yu.; Maksimov, E. E.; Alakhov, Yu. B.
ΑU
     Pushchino Branch, Shemyakin-Ovchinnikov Institute of Bioorganic Chemistry,
CS
     Russian Academy of Sciences, Pushchino, 142292, Russia
     Bioorganicheskaya Khimiya (1998), 24(7), 539-543
SO
     CODEN: BIKHD7; ISSN: 0132-3423
PB
     MAIK Nauka
     Journal
ÐΤ
     Russian
LΑ
CC
     9-11 (Biochemical Methods)
     Here we describe a method for obtaining a ready-to-use stabilized reaction
AΒ
     mixt. for in vitro translation of mRNA. We also demonstrate the
     stabilization of a complete translation mixt. contg. wheat germ ext.,
     amino acids, ATP, GTP, creatine phosphate, creatine kinase, and the
     reaction buffer by lyophilization in the presence of various sugars.
     greatest stabilizing effect is achieved by supplementing the mixt. with
     10% (mass/vol.) trehalose, which is also a unique translation
     activator, enhancing the translation of various mRNAs. A lyophilized
     complete translation mixt. contg. trehalose can be stored at
     4-80C for several months without losing its activity. The mixt. can be
     easily reconstituted by adding an aq. mRNA soln. and retains the potential
     for reproducible functioning. This allows the employment of such a
     cell-free translation system for anal. screening of a broad spectrum of
     compds. inhibiting translation at various stages.
     translation mRNA in vitro reaction mixt lyophilization preservation
ST
ΙT
     Freeze drying
       Preservation
     Storage
     Translation, genetic
        (a stabilized reaction mixt. for in vitro translation of mRNA)
IT
     RL: BPR (Biological process); BSU (Biological study, unclassified); PRP
     (Properties); BIOL (Biological study); PROC (Process)
        (a stabilized reaction mixt. for in vitro translation of mRNA)
     Amino acids, biological studies
ΙT
     Carbohydrates, biological studies
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (a stabilized reaction mixt. for in vitro translation of mRNA)
ΙT
     Wheat
        (germ, ext.; a stabilized reaction mixt. for in vitro translation of
                                           67-07-2, Creatine phosphate
     56-65-5, 5'-ATP, biological studies
ΙT
     86-01-1, 5'-GTP 99-20-7, Trehalose
                                          9001-15-4,
     Creatine kinase
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (a stabilized reaction mixt. for in vitro translation of mRNA)
ΙT
     99-20-7, Trehalose
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (a stabilized reaction mixt. for in vitro translation of mRNA)
     99-20-7 HCAPLUS
RN
     .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)
CN
Absolute stereochemistry. Rotation (+).
```

L72 ANSWER 17 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN

AN 1998:676219 HCAPLUS

DN 130:11753

TI Protein inactivation in amorphous sucrose and trehalose matrixes: effects of phase separation and crystallization

AU Sun, Wendell Q.; Davidson, Paul

CS School of Biological Sciences, National University of Singapore, Singapore, 119260, Singapore

SO Biochimica et Biophysica Acta (1998), 1425(1), 235-244 CODEN: BBACAQ; ISSN: 0006-3002

PB Elsevier Science B.V.

DT Journal

LA English

CC 6-3 (General Biochemistry)
 Section cross-reference(s): 7, 9

Trehalose is the most effective carbohydrate in preserving the AB structure and function of biol. systems during dehydration and subsequent storage. We have studied the kinetics of protein inactivation in amorphous glucose/sucrose (1:10, wt./wt.) and glucose/trehalose (1:10, wt./wt.) systems, and examd. the relationship between protein preservation, phase sepn. and crystn. during dry storage. The glucose/ trehalose system preserved glucose-6-phosphate dehydrogenase better than did the glucose/sucrose system with the same glass transition temp. (Tg). The Williams-Landel-Ferry kinetic anal. indicated that the superiority of the glucose/trehalose system over the glucose/sucrose system was possibly assocd. with a low free vol. and a low free vol. expansion at temps. above the Tg. Phase sepn. and crystn. during storage were studied using differential scanning calorimetry, and three sep. domains were identified in stored samples (i.e., sugar crystals, glucose-rich and disaccharide-rich amorphous domains). sepn. and crystn. were significantly retarded in the glucose/ trehalose system. Our data suggest that the superior stability of the trehalose system is assocd. with several properties of the trehalose glass, including low free vol., restricted mol. mobility and the ability to resist phase sepn. and crystn. during storage. protein preservation dehydration sucrose trehalose glass; phase ST sepn crystn protein storage carbohydrate

IT Enzyme kinetics

(of inhibition; protein inactivation in amorphous sucrose and trehalose matrixes with regard to effects of phase sepn. and crystn.)

IT Crystallization

Dehydration

Dehydration, physiological

Glass structure

Glass transition temperature

Phase separation

(protein inactivation in amorphous sucrose and trehalose matrixes with regard to effects of phase sepn. and crystn.)

```
IT
     Enzymes, biological studies
     Proteins, general, biological studies
     RL: BAC (Biological activity or effector, except adverse); BPR (Biological
     process); BSU (Biological study, unclassified); PEP (Physical, engineering
     or chemical process); BIOL (Biological study); PROC (Process)
        (protein inactivation in amorphous sucrose and trehalose
        matrixes with regard to effects of phase sepn. and crystn.)
ΙT
     Carbohydrates, biological studies
     RL: BPR (Biological process); BSU (Biological study, unclassified); PEP
     (Physical, engineering or chemical process); PRP (Properties); BIOL
     (Biological study); PROC (Process)
        (protein inactivation in amorphous sucrose and trehalose
        matrixes with regard to effects of phase sepn. and crystn.)
ΙT
     Preservation
        (protein; protein inactivation in amorphous sucrose and
        trehalose matrixes with regard to effects of phase sepn. and
ΙT
     9001-40-5, Glucose-6-phosphate dehydrogenase
     RL: BAC (Biological activity or effector, except adverse); BPR (Biological
    process); BSU (Biological study, unclassified); PEP (Physical, engineering
     or chemical process); BIOL (Biological study); PROC (Process)
        (protein inactivation in amorphous sucrose and trehalose
        matrixes with regard to effects of phase sepn. and crystn.)
ΙT
     50-99-7, Glucose, biological studies 57-50-1, Sucrose, biological
     studies 99-20-7, Trehalose
     RL: BPR (Biological process); BSU (Biological study, unclassified); PEP
     (Physical, engineering or chemical process); PRP (Properties); BIOL
     (Biological study); PROC (Process)
        (protein inactivation in amorphous sucrose and trehalose
        matrixes with regard to effects of phase sepn. and crystn.)
              THERE ARE 39 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
RE
(1) Aldous, B; Cryo-Letters 1995, V16, P181 HCAPLUS
(2) Arakawa, T; Adv Drug Deliv Rev 1993, V10, P1 HCAPLUS
(3) Cardona, S; J Food Sci 1997, V62, P105 HCAPLUS
(4) Carpenter, J; Biochemistry 1989, V28, P3916 HCAPLUS
(5) Chang, B; Arch Biochem Biophys 1996, V331, P249 HCAPLUS
(6) Crowe, J; Arch Biochem Biophys. 1983, V220, P477 HCAPLUS
(7) Crowe, J; Cryobiology 1994, V31, P355 MEDLINE
(8) Crowe, J; Water and Life 1992, P87
(9) Crowe, L; Biochim Biophys Acta 1984, V769, P141 HCAPLUS
(10) Crowe, L; Biophys J 1996, V71, P2087 HCAPLUS
(11) Ding, S; J Therm Anal 1996, V47, P1391 HCAPLUS
(12) Duddu, S; Pharmaceut Res 1997, V14, P591 HCAPLUS
(13) Franks, F; Trends Biotechnol 1994, V12, P114 HCAPLUS
(14) Gaffney, S; J Chem Soc Faraday Trans 1988, V84, P2545 HCAPLUS
(15) Green, J; J Phys Chem 1989, V93, P2880 HCAPLUS
(16) Koster, K; J Therm Anal 1996, V47, P1581 HCAPLUS
(17) Leopold, A; Stress Responses in Plants: Adaptation and Acclimation
   Mechanisms 1990, P37
(18) Leslie, S; Appl Environ Microbiol 1995, V61, P3592 HCAPLUS
(19) Mazzobre, M; Food Sci Technol 1997, V30, P324 HCAPLUS
(20) Miller, D; Pharmaceut Res 1997, V14, P578 HCAPLUS
(21) Mouradian, R; Biochim Biophys Acta 1984, V778, P615 HCAPLUS
(22) O'Brien, J; J Food Sci 1996, V61, P679 HCAPLUS
(23) Roser, B; BioPharm 1991, V4, P47 HCAPLUS
(24) Saleki-Gerhardt, A; J Pharmaceut Sci 1995, V84, P318 HCAPLUS
(25) Sastry, G; Biochemistry 1997, V36, P7097 HCAPLUS
(26) Schebor, C; J Food Eng 1996, V30, P269
(27) Slade, L; Crit Rev Food Sci Nutr 1991, V30, P115 HCAPLUS
(28) Sun, W; Ann Bot 1994, V74, P601
(29) Sun, W; Ann Bot 1997, V79, P291
```

(30) Sun, W; Biophys J 1996, V70, P1769 HCAPLUS

```
(31) Sun, W; Comp Biochem Physiol 1997, V117A, P327 HCAPLUS
```

(32) Sun, W; Cryo-Letters 1997, V18, P99 HCAPLUS

(33) Sun, W; Cryo-Letters 1998, V19, P105

(34) Sun, W; Cryo-Letters 1998, V19, P66 HCAPLUS

(35) Sun, W; Physiol Plant 1994, V90, P621 HCAPLUS

(36) Uritani, M; J Biochem 1995, V117, P774 HCAPLUS

(37) Williams, M; J Am Chem Soc 1955, V77, P3701 HCAPLUS

(38) Wolkers, W; Biochim Biophys Acta 1998, V1379, P83 HCAPLUS

(39) Yoshioka, S; Pharmaceut Res 1997, V14, P736 HCAPLUS

IT 99-20-7, Trehalose

RL: BPR (Biological process); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); PRP (Properties); BIOL (Biological study); PROC (Process)

(protein inactivation in amorphous sucrose and trehalose matrixes with regard to effects of phase sepn. and crystn.)

RN 99-20-7 HCAPLUS

CN ..alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

L72 ANSWER 18 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN

AN 1995:344703 HCAPLUS

DN 122:155595

TI Do trehalose and dimethyl sulfoxide affect intermembrane forces?

AU Pincet, Frederic; Perez, Eric; Wolfe, Joe

CS Laboratoire de Physique Statistique, Ecole Normale Superieure, Paris, 75231, Fr.

SO Cryobiology (1994), 31(6), 531-9 CODEN: CRYBAS; ISSN: 0011-2240

DT Journal

LA English

CC 9-11 (Biochemical Methods)
 Section cross-reference(s): 6

The sugar trehalose is produced in some organisms that survive AΒ dehydration and desiccation, and it preserves the integrity of membranes in model systems exposed to dehydration and freezing. DMSO, a solute which permeates membranes, is added to cell suspensions in many protocols for cryopreservation. Using a surface forces app., the authors measured the very large, short-range repulsion between phosphatidylcholine bilayers in water and in solns. of trehalose, sorbitol, and DMSO. To the resoln. of the technique, the force-distance curves between bilayers are unchanged by the addn. of trehalose or sorbitol in concns. >1 kmol/m3. A relatively small increase in adhesion in the presence of trehalose and sorbitol solns. may be explained by their osmotic effects. The partitioning of trehalose between aq. solns. and lamellar phases of dioleoylphosphatidylcholine was measured gravimetrically. The amt. of trehalose that preferentially adsorbs near membrane surfaces is at most small. The presence of DMSO in water (1:2 by vol.) makes very little difference to the short-range interaction between deposited bilayers, but it sometimes perturbs them in

ways that vary among expts.: free bilayers and/or fusion of the deposited bilayers were each obsd. in about one-third of the expts.

ST phospholipid membrane dehydration freezing protectant; cryoprotectant phospholipid bilayer intermembrane force; trehalose intermembrane force bilayer membrane; DMSO intermembrane force bilayer membrane

IT Organ preservation

(cryopreservation; trehalose and DMSO effects on intermembrane forces study in phospholipid bilayers)

IT Cryoprotectants

Dehydration, biological

Freezing

(trehalose and DMSO effects on intermembrane forces study in phospholipid bilayers)

IT Membrane, biological

(bilayer, trehalose and DMSO effects on intermembrane forces study in phospholipid bilayers)

IT 68737-67-7, Dioleoylphosphatidylcholine

RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); BIOL (Biological study); OCCU (Occurrence); PROC (Process)

(trehalose and DMSO effects on intermembrane forces study in phospholipid bilayers)

IT 50-70-4, Sorbitol, biological studies 67-68-5, DMSO, biological studies 99-20-7, Trehalose

RL: BUU (Biological use, unclassified); PEP (Physical, engineering or chemical process); BIOL (Biological study); PROC (Process); USES (Uses) (trehalose and DMSO effects on intermembrane forces study in phospholipid bilayers)

IT 99-20-7, Trehalose

RL: BUU (Biological use, unclassified); PEP (Physical, engineering or chemical process); BIOL (Biological study); PROC (Process); USES (Uses) (trehalose and DMSO effects on intermembrane forces study in phospholipid bilayers)

RN 99-20-7 HCAPLUS

CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

L72 ANSWER 19 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN

AN 1992:169671 HCAPLUS

DN 116:169671

TI Method and composition for the preservation of red blood cells by lyophilization

IN Rudolph, Alan S.; Larry, Joseph Pat

PA United States Dept. of the Navy, USA

SO U. S. Pat. Appl., 17 pp. Avail. NTIS Order No. PAT-APPL. 7-659 765. CODEN: XAXXAV

DT Patent

LA English

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CC
     9-11 (Biochemical Methods)
     Section cross-reference(s): 13
FAN.CNT 1
                      KIND DATE
     PATENT NO.
                                           APPLICATION NO.
                                                            DATE
                           _____
                                           -----
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                      A0
                            19920201
                                           US 1991-659765
                                                            19910225 <--
PΙ
     US 659765
                            19930907
     US 5242792
                       Α
PRAI US 1991-659765
                            19910225
                                      <--
     The title compn. includes a permeabilizing agent (e.g. glycerol), a
     preserving agent, and a buffered solvent. The compn. is used to prep.
     erythrocytes for lyophilization, as well as to rehydrate them following
     lyophilization. A method using the compn. for erythrocyte lyophilization
     and rehydration is described. Trehalose is the preferred
     protective agent. When glycerol and trehalose were used as
     permeabilizing and protective agents, resp., the optimum recovery of oxyHb
     was obsd. with 10% glycerol and 500 mM trehalose, where there
     was no measurable metHb; these conditions also resulted in the highest
     recovery of the P50 value (an expression of O-carrying capacity).
     erythrocyte lyophilization rehydration soln; trehalose glycerol
ST
     erythrocyte lyophilization rehydration
ΙT
     Preservatives
        (in erythrocyte lyophilization and rehydration compn.)
ΙT
     Erythrocyte
        (lyophilization and rehydration of, compn. for)
ΙT
     Freeze drying
        (of erythrocyte, compn. for)
TT
     Hemoglobins, met-
     Hemoglobins, oxy-
     RL: BIOL (Biological study)
        (trehalose-contg. erythrocyte lyophilization and rehydration
        compn. effect on)
ΙT
     Biological transport
        (permeation, agent enhancing, in erythrocyte lyophilization and
        rehydration compn.)
ΙT
     Hydration, biological
        (re-, of erythrocyte, compn. for)
TT
     7782-44-7, Oxygen, biological studies
     RL: BIOL (Biological study)
        (carrying capacity, trehalose-contg. erythrocyte
        lyophilization and rehydration compn. effect on)
     56-81-5, Glycerol, biological studies 99-20-7, Trehalose
TΤ
     RL: BIOL (Biological study)
        (in erythrocyte lyophilization and rehydration compn.)
IT
     99-20-7, Trehalose
     RL: BIOL (Biological study)
        (in erythrocyte lyophilization and rehydration compn.)
RN
     99-20-7 HCAPLUS
     .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI)
CN
                                                                (CA INDEX NAME)
Absolute stereochemistry. Rotation (+).
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L72 ANSWER 20 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN
AN
    1992:124367 HCAPLUS
DN
     116:124367
     Stabilization of biological macromolecular substances and other organic
TΙ
     compounds with nonreducing polyhydroxy glycosides or oligosaccharides
IN
     Roser, Bruce Joseph; Colaco, Camilo
PΑ
     Quadrant Holdings Cambridge Ltd., UK
SO
     PCT Int. Appl., 24 pp.
    CODEN: PIXXD2
DT
    Patent
LA
    English
     ICM C12N009-96
IC
     ICS A61K047-26
CC
     9-2 (Biochemical Methods)
FAN.CNT 1
     PATENT NO.
                     KIND DATE
                                          APPLICATION NO. DATE
                     ----
                                         ______
     -----
                                        WO 1991-GB759 19910514 <--
                     A1 19911128
    WO 9118091
PΙ
        W: AT, AU, BB, BG, BR, CA, CH, DE, DK, ES, FI, GB, HU, JP, KP, KR,
            LK, LU, MC, MG, MW, NL, NO, PL, RO, SD, SE, SU, US
         RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, DE, DK, ES, FR, GA, GB, GR,
            IT, LU, ML, MR, NL, SE, SN, TD, TG
                                          AU 1991-78725
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    EP 541556
                      В1
                           19980916
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE
                    T2 19931125
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    JP 3101320
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                     T3 19990301
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                                          US 1994-255565
                                                           19940608 <--
    US 5621094
PRAI GB 1990-10742
                     Α
                           19900514 <--
                          19910514 <--
    WO 1991-GB759
                     Α
                          19921214 <--
    US 1992-965384
                     В1
     (Bio)org. compds. are preserved in a dry state, at elevated temps., and/or
AB
     under irradn. with nonreducing oligosaccharides or polyhydroxy glycosides.
    Restriction endonuclease PstI was dried at room temp. in the presence of
     trehalose then stored for 2 wks at 37.degree.. The enzyme
     retained 100% of its original activity after this treatment.
    bioorg compd preservation drying temp; irradn bioorg compd preservation;
ST
    oligosaccharide nonreducing bioorg compd preservation; polyhydroxy
    glycoside nonreducing preservation biomol
IT
    Galactosides
    Glycosides
    Oligosaccharides
    RL: ANST (Analytical study)
        (nonreducing, bioorg. compd. stabilization to drying and elevated temp.
        and irradn. with)
IT
    Preservation
        (of org. and bioorg. compds., to drying and elevated temp. and irradn.,
        with nonreducing polyhydroxy glycosides and oligosaccharides)
IΤ
    Temperature effects, biological
        (on org. and bioorg. compds., stabilization with nonreducing
       polyhydroxy glycosides and oligosaccharides in relation to.)
ΙT
    Drying
        (stabilization of org. and bioorg. compds. to, with nonreducing
        polyhydroxy glycosides and oligosaccharides)
TT
    Organic compounds, miscellaneous
    RL: MSC (Miscellaneous)
        (stabilization of, to drying and elevated temp. and irradn., with
        nonreducing polyhydroxy glycosides and oligosaccharides)
IT
    Light stabilizers
```

(UV, nonreducing polyhydroxy glycosides and oligosaccharides as, for org. and bioorg. compds.)

IT Carbohydrates and Sugars, uses

RL: USES (Uses)

(alditols, nonreducing, glycosides, bioorg. compd. stabilization to drying and elevated temp. and irradn. with)

IT Organic compounds, miscellaneous

RL: MSC (Miscellaneous)

(biol., stabilization of, to drying and elevated temp. and irradn., with nonreducing polyhydroxy glycosides and oligosaccharides)

IT Oligosaccharides

RL: ANST (Analytical study)

(di-, nonreducing, bioorg. compd. stabilization to drying and elevated temp. and irradn. with)

IT Alcohols, uses

RL: USES (Uses)

(polyhydric, nonreducing, glycosides, bioorg. compd. stabilization to drying and elevated temp. and irradn. with)

IT Phycoerythrins

RL: ANST (Analytical study)

(R-, stabilization to drying of, nonreducing oligosaccharides and polyhydroxy glycosides in)

IT 13718-94-0, Isomaltulose 64519-82-0, Palatinit 50-70-4, Sorbitol, biological studies 57-50-1, Sucrose, biological studies 69-65-8, Mannitol 99-20-7, Trehalose 470-55-3, Stachyose

512-69-6, Raffinose 534-73-6 585-86-4, Lactitol 585-88-6, Maltitol 597-12-6, Melezitose 4233-70-9

RL: ANST (Analytical study)

(bioorg. compd. stabilization to drying and elevated temp. and irradn. with)

IT 9012-36-6P, Agarose

RL: PREP (Preparation)

(gels, stabilization to drying of, nonreducing oligosaccharides and polyhydroxy glycosides in)

IT 9003-99-0D, Peroxidase, fusion products with Ig F(ab)2 fragment 81295-32-1, Restriction endonuclease PstI

RL: ANST (Analytical study)

(stabilization to drying of, nonreducing oligosaccharides and polyhydroxy glycosides in)

IT 99-20-7, Trehalose

RL: BIOL (Biological study)

(bioorg. compd. stabilization to drying and elevated temp. and irradn. with)

RN 99-20-7 HCAPLUS

CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

L72 ANSWER 21 OF 21 HCAPLUS COPYRIGHT 2003 ACS on STN AN 1988:507400 HCAPLUS

```
109:107400
DN
ΤI
     Polyhydroxy compounds in the preservation of biological materials such as
     hemoglobin, erythrocytes, liposomes, and cells
ΙN
     Hayward, James Arthur; Johnston, David Samuel
PA
     Biocompatibles Ltd., UK
SO
     PCT Int. Appl., 24 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
IC
     ICM C07K003-00
     ICS C07K013-00; A61K009-50; A61K035-48
    A61K037-04; C12N009-36; C12N009-96
     9-11 (Biochemical Methods)
FAN.CNT 1
     PATENT NO.
                      KIND DATE
                                           APPLICATION NO.
                                                            DATE
                      <del>-</del>---
                                           -----
                                                            -----
     WO 8705300
                      A2
                            19870911
                                           WO 1987-GB143
                                                            19870227 <--
     WO 8705300
                      A3
                            19871022
        W: JP, US
         RW: AT, BE, CH, DE, FR, GB, IT, LU, NL, SE
                      A1
                           19880316
                                          EP 1987-901561
                                                            19870227 <--
        R: AT, BE, CH, DE, FR, GB, IT, LI, LU, NL, SE
                                          JP 1987-501509
     JP 63502592
                       T2
                            19880929
                                                            19870227 <--
                                     <--
PRAI GB 1986-4983
                            19860228
                            19870227 <--
     WO 1987-GB143
     A material having a water-dependent structure is preserved by contacting
AB
     the material with an aq. soln. of a polyhydroxy compd. and then removing
     water. Carbohydrates were added to aq. Hb solns. (0.015M) to 0.25M concn.
     and the mixts. were dehydrated. The percentage of metHb formed was
     measured in rehydrated protein. In the absence of sugar, the percentage
     increased to 90% of the total. Each of the carbohydrates tested
     (arabinose, galactose, fucose, glucose, mannose, maltose, lactose,
     trehalose, sucrose) protected the Hb against oxidative damage.
     This protective effect was manifest even in samples that had been stored
     in the dried state for >3 mo.
     polyol biol material preservation; Hb preservation carbohydrate;
ST
     erythrocyte preservation carbohydrate; liposome preservation carbohydrate;
     cell preservation carbohydrate
ΙT
     Hexoses
     Monosaccharides
     Oligosaccharides
     Pentoses
     Carbohydrates and Sugars, biological studies
     RL: ANST (Analytical study)
        (in biol. materials preservation)
ΙT
     Dehydration, biological
        (of biol. materials, polyhydroxy compds. in)
IT
     Blood preservation
        (polyhydroxy compds. in)
IT
     Biological materials
     Cell
     Liposome
     Proteins, biological studies
     Hemoglobins
     Hemoglobins, oxy-
     RL: BIOL (Biological study)
        (preservation of, polyhydroxy compds. in)
ΙT
     Oligosaccharides
     RL: ANST (Analytical study)
        (di-, in biol. materials preservation)
ΙT
     Organelle
        (hemosome, preservation of, polyhydroxy compds. in)
     Carbohydrates and Sugars, biological studies
IT
```

RL: BIOL (Biological study) (nonreducing, in biol. materials preservation) IT Hydroxy compounds RL: ANST (Analytical study) (poly-, in biol. materials preservation) Alcohols, biological studies IT RL: BIOL (Biological study) (polyhydric, in biol. materials preservation) Carbohydrates and Sugars, biological studies ΙT RL: BIOL (Biological study) (trioses, in biol. materials preservation) 63-42-3, Lactose 69-79-4, Maltose 147-81-9, Arabinose 2438-80-4, ΙT 57-50-1, Sucrose, biological studies 3458-28**-4**, Mannose RL: ANST (Analytical study) (in Hb preservation) 99-20-7, Trehalose 50-99-7, Glucose, biological IT studies 59-23-4, Galactose, biological studies RL: ANST (Analytical study) (in biol. materials preservation) ΙT 9001-63-2, Lysozyme RL: PROC (Process) (preservation of, polyhydroxy compds. in) IT 99-20-7, Trehalose RL: ANST (Analytical study) (in biol. materials preservation) 99-20-7 HCAPLUS RN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

CN

=> d 173 all hitstr tot

ANSWER 1 OF 11 HCAPLUS COPYRIGHT 2003 ACS on STN L73 2003:656142 HCAPLUS ANDN Bulk drying and the effects of inducing bubble nucleation ΤI Bronshtein, Victor; Bracken, Kevin R.; Campbell, John G. ΙN PA U.S. Pat. Appl. Publ., 22 pp. SO CODEN: USXXCO DT Patent LA English ICM B01J013-02 ICS B01J013-04 NCL 264004100; 264004600 9-16 (Biochemical Methods) FAN.CNT 1 DATE KIND DATE APPLICATION NO. PATENT NO.

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20030821
                                           US 2002-274719
                                                             20021018
     US 2003155669
                       A1
PRAI US 2001-345322P
                       Р
                            20011019
     The present invention discloses app. and methods of inducing bubble
     nucleation to overcome problems commonly assocd. with preservation by foam
     formation. Specifically, the invention relates to methods of using bubble
     nucleation in foam formation to preserve sensitive biol. materials.
     Preferred methods of inducing bubble nucleation include, mixing, chamber
     rotation, crystals, and ultrasound.
     bulk drying bubble nucleation
ST
     Convective flow
IT
        (Forced; bulk drying and the effects of inducing bubble nucleation)
IT
     Mixers (processing apparatus)
        (Magnetic; bulk drying and the effects of inducing bubble nucleation)
IT
     Containers
        (Process; bulk drying and the effects of inducing bubble nucleation)
IT
     Pressure
        (Vacuum; bulk drying and the effects of inducing bubble nucleation)
IT
     Apparatus
     Biological materials
     Blades
     Boiling
     Bubbles
     Coating materials
     Containers
     Crystals
       Drying
     Foaming
     Freezing
     Frequency
     Heating
     Mixing
     Nucleation
       Preservation
     Pressure
     Pressure sensors
     Rings (apparatus)
     Rotation
     Shear
     Solutions
     Sound and Ultrasound
     Suspensions
     Temperature
     Temperature sensors
        (bulk drying and the effects of inducing bubble nucleation)
IT
     Amino acids, biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (bulk drying and the effects of inducing bubble nucleation)
ΙT
     Fluoropolymers, uses
     RL: TEM (Technical or engineered material use); USES (Uses)
        (bulk drying and the effects of inducing bubble nucleation)
IT
     Vacuum
        (pressure; bulk drying and the effects of inducing bubble nucleation)
ΙT
     Mixers (processing apparatus)
        (stirrers, Bars; bulk drying and the effects of inducing bubble
        nucleation)
     50-99-7, Glucose, biological studies
                                             57-48-7, D-Fructose, biological
IT
               57-50-1, Sucrose, biological studies
                                                       58-08-2, Caffeine,
                          87-89-8, Inositol 99-20-7,
     biological studies
     Trehalose
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (bulk drying and the effects of inducing bubble nucleation)
```

IT 9002-84-0, Teflon

RL: TEM (Technical or engineered material use); USES (Uses) (bulk drying and the effects of inducing bubble nucleation)

IT 99-20-7, Trehalose

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(bulk drying and the effects of inducing bubble nucleation)

RN 99-20-7 HCAPLUS

CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+)'.

L73 ANSWER 2 OF 11 HCAPLUS COPYRIGHT 2003 ACS on STN

AN 2003:126473 HCAPLUS

DN 139:115030

TI Stabilization of membranes in human platelets freeze-dried with trehalose

AU Crowe, John H.; Tablin, Fern; Wolkers, Willem F.; Gousset, Karine; Tsvetkova, Nelly M.; Ricker, Josette

CS Center for Biostabilization, University of California, Davis, CA, 95616, USA

SO Chemistry and Physics of Lipids (2003), 122(1-2), 41-52 CODEN: CPLIA4; ISSN: 0009-3084

PB Elsevier Science Ltd.

DT Journal

LA English

CC 13-5 (Mammalian Biochemistry)
 Section cross-reference(s): 9

Human blood platelets are normally stored in blood banks for 3-5 days, after which they are discarded. We have launched an effort at developing means for preserving the platelets for long term storage. In previous studies we have shown that trehalose can be used to preserve biol. membranes and proteins during drying and have provided evidence concerning the mechanism. A myth has grown up about special properties of trehalose, which we discuss here and clarify some of what is fact and what is misconception. We have found a simple way of introducing this sugar into the cytoplasm of platelets and have successfully freeze-dried the trehalose-loaded platelets, with very promising results. We present evidence that membrane microdomains are maintained intact in the platelets freeze-dried with trehalose. Finally, we propose a possible mechanism by which the microdomains are preserved.

ST **trehalose** platelet preservation membrane stabilization freeze drying

IT Blood preservation

Cell membrane

Freeze drying

Human

Platelet (blood)

(stabilization of membranes in human platelets freeze-dried with trehalose)

lucas - 10 / 018834 99-20-7, Trehalose TT RL: BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses) (stabilization of membranes in human platelets freeze-dried with RE.CNT 59 THERE ARE 59 CITED REFERENCES AVAILABLE FOR THIS RECORD (1) Aldous, B; Cryo-Lett 1995, V16, P181 HCAPLUS (2) Arguelles, J; Arch Microbiol 2000, V174, P217 HCAPLUS (3) Beattie, G; Diabetes 1997, V46, P519 HCAPLUS (4) Benaroudj, N; J Biol Chem 2001, V26, P24261 (5) Branca, C; J Chem Phys 1999, V111, P281 HCAPLUS (6) Brown, D; Annu Rev Cell Dev Biol 1998, V14, P111 HCAPLUS (7) Brown, D; J Membr Biol 1998, V164, P103 HCAPLUS (8) Buera, M; Biotechnol Prog 1999, V15, P577 HCAPLUS (9) Chen, T; Cryobiology 2000, V40, P277 HCAPLUS (10) Chen, T; Cryobiology 2001, V43, P168 HCAPLUS (11) Crowe, J; Annu Rev Physiol 1998, V6, P73 (12) Crowe, J; Biochem J 1987, V242, P1 HCAPLUS (13) Crowe, J; Cryobiology 1997, V35, P20 HCAPLUS (14) Crowe, J; Cryobiology 2001, V43, P89 HCAPLUS (15) Crowe, J; Liposome Technology, second ed 1992 (16) Crowe, J; Proc Natl Acad Sci USA 1987, V84, P1537 HCAPLUS (17) Crowe, L; Biochim Biophys Acta 1988, V946, P193 HCAPLUS (18) Crowe, L; Biophys J 1996, V71, P2087 HCAPLUS (19) de Castro, A; FEBS Lett 2000, V487, P199 (20) Eroglu, A; Nat Biotechnol 2000, V18, P163 HCAPLUS (21) Esteves, M; Biotechnol Lett 2000, V22, P417 HCAPLUS (22) Gimeno-Alcaniz, J; Biotechnol Lett 1999, V21, P271 HCAPLUS (23) Gordon, S; Cryobiology 2001, V43, P182 HCAPLUS (24) Gousset, K; Biochemistry Submitted for publication 2003 (25) Gousset, K; J Cell Physiol 2002, V190, P117 HCAPLUS (26) Green, J; J Phys Chem 1989, V93, P2880 HCAPLUS (27) Guo, N; Nat Biotechnol 2000, V18, P168 HCAPLUS (28) Hays, L; Cryobiology 2001, V42, P88 HCAPLUS (29) Hays, L; Proc Natl Acad Sci 1995, V93, P6835 (30) Hincha, D; Eur J Biochem 2000, V267, P535 HCAPLUS (31) Hoekstra, F; Comp Biochem Physiol (A) 1997, V117A, P335 HCAPLUS (32) Joo, W; Biotechnol Lett 2000, V22, P1021 HCAPLUS (33) Kitahara, A; Br J Plast Surg 1998, V51, P118 MEDLINE (34) Koster, K; Biophys J 2000, V78, P1932 HCAPLUS (35) Leslie, S; Econ Environ Microbiol 1995, V61, P3592 HCAPLUS (36) Leyman, B; Trends Plant Sci 2001, V6, P510 HCAPLUS (37) Li, S; J Pharm Sci 1996, V85, P873 HCAPLUS (38) Liu, Q; J Am Chem Soc 1997, V119, P7851 HCAPLUS

- (39) Mansure, J; Biochim Biophys Acta Bio-Membr 1994, V1191, P309 HCAPLUS
- (40) Matsuo, T; Br J Ophthalmol 2001, V85, P610 MEDLINE
- (41) Mendelsohn, R; Chem Phys Lipids 1998, V96, P141 HCAPLUS
- (42) Mendelsohn, R; Chem Phys Lipids 1998, V96, P141 HCAPLUS
- (43) Nishizaki, Y; Nutr Res 2000, V20, P653 HCAPLUS
- (44) Norcia, M; Compositions and methods for wound management 2000, P424
- (45) Otsubo, M; Hort Sci 2000, V35, P1107 HCAPLUS
- (46) O'Brien, J; J Food Sci 1996, V61, P679 HCAPLUS
- (47) Potts, M; Trends Microbiol 2001, V9, P553 HCAPLUS
- (48) Ricker, J; Biophysical Journal In press 2002
- (49) Schebor, C; Biotechnol Prog 1997, V13, P857 HCAPLUS (50) Schebor, C; Biotechnol Prog 2000, V16, P163 HCAPLUS
- (51) Schebor, C; Lebensm-Wiss Technol 1999, V32, P481 HCAPLUS
- (52) Sun, W; Biophys J 1996, V70, P1769 HCAPLUS
- (53) Sussich, F; Carbohydr Res 2001, V334, P165 HCAPLUS
- (54) Tablin, F; Cryobiology 2001, V43, P114 HCAPLUS
- (55) Tablin, F; J Cell Physiol 1996, V168, P305 HCAPLUS
- (56) Tsvetkova, N; Mol Membr Biol 1999, V16, P265 HCAPLUS

(57) Wolkers, W; Cryobiology 2001, V42, P79 HCAPLUS

(58) Wolkers, W; Mol Membrane Biology in press 2003

(59) Yoshizane, C; Nutr Res 2000, V20, P1485 HCAPLUS

IT 99-20-7, Trehalose

RL: BSU (Biological study, unclassified); BUU (Biological use,

unclassified); BIOL (Biological study); USES (Uses)

(stabilization of membranes in human platelets freeze-dried with

trehalose)

RN 99-20-7 HCAPLUS

CN .alpha.-D-Glucopyranoside; .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

L73 ANSWER 3 OF 11 HCAPLUS COPYRIGHT 2003 ACS on STN

AN 2003:58684 HCAPLUS

DN 138:119601

TI Vacuum-mediated desiccation protection of cells

IN Levine, Fred

PA Regents of the University of California, USA

SO U.S. Pat. Appl. Publ., 29 pp. CODEN: USXXCO

DT Patent

LA English

IC ICM A01N001-02 ICS C12N005-06

NCL 435002000; 435325000; 514053000

CC 9-11 (Biochemical Methods)
 Section cross-reference(s): 3

FAN.CNT 1

FAN.	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	US 2003017444	A1 ·	20030123	US 2001-812042	20010319
	US 6528309	B2	20030304		
דעממ	rrs 2001-812042		20010319		

AB The present invention provides methods and compns. for the protection and storage of cells. In particular, the present invention provides methods and compns. for the vacuum-mediated desiccation protection of mammalian cells. In particularly preferred embodiments, cells are treated with a carbohydrate (e.g., a disaccharide) prior to vacuum-mediated desiccation.

ST vacuum desiccation human cell storage carbohydrate trehalose expression preservation

IT Animal cell line

(12F; vacuum-mediated desiccation protection of cells)

IT Adenoviral vectors

(expressing otsA and otsB; vacuum-mediated desiccation protection of cells)

IT Animal cell

(mammalian; vacuum-mediated desiccation protection of cells)

IT Alcohols, biological studies

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES

(Uses)

(polyhydric; vacuum-mediated desiccation protection of cells)

IT Adhesion, biological

Animal cell

Drying

Genetic methods

Human

Preservation

Storage

Thermal shock

Vacuum

(vacuum-mediated desiccation protection of cells)

IT Carbohydrates, biological studies

Disaccharides

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(vacuum-mediated desiccation protection of cells)

IT 99-20-7, Trehalose

RL: BSU (Biological study, unclassified); BUU (Biological use,

unclassified); BIOL (Biological study); USES (Uses)

(expression of; vacuum-mediated desiccation protection of cells)

IT 7782-44-7, Oxygen, biological studies

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(less that 3%; vacuum-mediated desiccation protection of cells)

IT 488762-68-1 488762-69-2 488762-70-5 488762-71-6

RL: PRP (Properties)

(unclaimed nucleotide sequence; vacuum-mediated desiccation protection of cells)

IT 99-20-7, Trehalose

RL: BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(expression of; vacuum-mediated desiccation protection of cells)

RN 99-20-7 HCAPLUS

CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

L73 ANSWER 4 OF 11 HCAPLUS COPYRIGHT 2003 ACS on STN

AN 2002:690028 HCAPLUS

DN 138:398272

TI The glass transition temperature of mixtures of trehalose and hydroxyethyl starch

AU Chen, Tani; Bhowmick, Sankha; Sputtek, Andreas; Fowler, Alex; Toner, Mehmet

CS Massachusetts General Hospital, Center for Engineering in Medicine and Surgical Services, Harvard Medical School and Shriners Hospital for Children, Boston, MA, 02114, USA

SO Cryobiology (2002), 44(3), 301-306 CODEN: CRYBAS; ISSN: 0011-2240

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PB
     Elsevier Science
\mathsf{DT}
     Journal
     English
LĄ
CC
     9-11 (Biochemical Methods)
     Although mixts. of HES and sugars are used to preserve cells during
AΒ
     freezing or drying, little is known about the glass transition of HES, or
     how mixts. of HES and sugars vitrify. These difficulties may be due to
     the polydispersity between HES samples or differences in prepn.
     techniques, as well as problems in measuring the glass transition temp.
     (Tg) using differential scanning calorimetry (DSC). In this report, we
     examine the Tq of mixts. of HES and trehalose sugar with <1%
     moisture content using DSC measurements. By extrapolating these
     measurements to pure HES using the Gordon-Taylor and Fox equations, we
     were able to est. the Tg of our HES sample at 44 .degree.C. These results
     were addnl. confirmed by using mixts. of glucose-HES which yielded a
     similar extrapolated Tg value. Our approach to estq. the glass transition
     temp. of HES may be useful in other cases where glass transitions are not
     easily identified.
ST
     glass transition temp trehalase hydroxyethyl starch cryopreservation;
     cryoprotectant glass transition temp modeling
ΙT
     Simulation and Modeling, physicochemical
        (Fox equation; glass transition temp. of mixts. of trehalose
        starch used in cell cryopreservation)
     Simulation and Modeling, physicochemical
ΙT
        (Gordon-Taylor equation; glass transition temp. of mixts. of
        trehalose starch used in cell cryopreservation)
     Cell
TΤ
       Cryopreservation
     Cryoprotectants
      Freeze drying
     Glass transition temperature
        (glass transition temp. of mixts. of trehalose starch used in
        cell cryopreservation)
ΙT
     99-20-7, Trehalose
                          9005-27-0, Hydroxyethyl starch
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (glass transition temp. of mixts. of trehalose starch used in
        cell cryopreservation)
RE.CNT
              THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD
(1) Banks, W; Br J Pharmacol 1973, V47, P172 HCAPLUS
(2) Chen, T; Cryobiology 2000, V40, P277 HCAPLUS
(3) Crowe, J; Annu Rev Physiol 1998, V60, P73 HCAPLUS
(4) Crowe, J; Biochem Biophys Acta 1996, V1280, P187 HCAPLUS
(5) Crowe, J; Cryobiology 1997, V35, P20 HCAPLUS
(6) Fox, T; J Appl Phys 1950, V21, P581 HCAPLUS
(7) Gordon, M; J Appl Chem 1952, V2, P493 HCAPLUS
(8) Jochem, M; Cryobiology 1987, V24, P513 HCAPLUS
(9) Koerber, C; Cryobiology 1980, V17, P54 HCAPLUS
(10) Kresin, M; Cryo-Lett 1992, V13, P371 HCAPLUS
(11) Rindler, V; Cryobiology 1999, V38, P2 HCAPLUS
(12) Roos, Y; Biotechnol Prog 1991, V7, P49 HCAPLUS
(13) Scheiwe, M; Cryobiology 1982, V19, P461 HCAPLUS
(14) Spieles, G; Cryo-Lett 1996, V17, P43
(15) Sputtek, A; Z Klin Med 1991, V46, P1567 HCAPLUS
(16) Wolfe, J; Cryobiology 1999, V39, P103 HCAPLUS
(17) Wunderlich, B; Thermal Analysis 1990
     99-20-7, Trehalose
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (glass transition temp. of mixts. of trehalose starch used in
        cell cryopreservation)
```

99-20-7 HCAPLUS

RN

CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

```
ANSWER 5 OF 11 HCAPLUS COPYRIGHT 2003 ACS on STN
     2002:638128 HCAPLUS
AN
DN
     137:152032
     Erythrocytic cells and method for preserving cells
ΤI
     Crowe, John H.; Crowe, Lois M.; Tablin, Fern; Wolkers, Willem F.;
IN
     Tsvetkova, Nelly M.; Oliver, Ann F.
PA
     U.S. Pat. Appl. Publ., 63 pp., Cont.-in-part of U.S. Ser. No. 927,760.
SO
     CODEN: USXXCO
DT
     Patent
LA
     English
IC
     ICM A61K048-00
     ICS A61K031-56
NCL
     424093210
     9-11 (Biochemical Methods)
     Section cross-reference(s): 13
FAN.CNT 3
     PATENT NO.
                      KIND
                            DATE
                                           APPLICATION NO.
                                                             DATE
                      ____
                            -----
                                           ______
                                                             _____
     US 2002114791
                       A1
                            20020822
                                           US 2002-52162
                                                             20020116
     US 2001019819
                       A1
                            20010906
                                           US 2001-828627
                                                             20010405
     US 2002076445
                       A1
                            20020620
                                           US 2001-927760
                                                             20010809
     WO 2003014331
                       A1
                            20030220
                                           WO 2002-US24773
                                                            20020805
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AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG PRAI US 2000-501773 В2 20000210 US 2001-828627 20010405 A2 US 2001-927760 .A2 20010809 US 2002-52162 Α 20020116

AB The invention concerns a dehydrated compn. is provided that includes freeze-dried erythrocytic cells. Alc. (e.g., sterol or cholesterol) is at least partially removed from erythrocytic cells including erythrocytic membranes. After removal of at least part of the alc., the erythrocytic cells have a low phase transition temp. range, an intermediate phase transition temp. range, and a high phase transition temp. range. The erythrocytic cells may be loaded with an oligosaccharide (e.g., trehalose) which preserves biol. properties during freeze-drying and rehydration. A process for increasing cooperativity of a phase

transition of an erythrocytic cell. A process for preserving and/or increasing the survival of dehydrated erythrocytic cells, including storing dehydrated erythrocytic cells having a residual water content equal to or less than about 0.30 g of water per g of dry wt. erythrocytic ST eukaryote cell culture temp preservation trehalose oligosaccharide phase transition ΙT Animal cell line (293H; erythrocytic cells and method for preserving cells) ITAnimal tissue culture Cryopreservation Cryoprotectants Dehydration, physiological Erythrocyte Freeze drying Heating Membrane, biological Phase transition Platelet (blood) Preservatives Temperature effects, biological Washing (erythrocytic cells and method for preserving cells) ΙT Oligosaccharides, biological studies RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses) (erythrocytic cells and method for preserving cells) IT Alcohols, properties Steroids, properties RL: PRP (Properties); REM (Removal or disposal); PROC (Process) (erythrocytic cells and method for preserving cells) ΙT (lipid phase; erythrocytic cells and method for preserving cells) IT Animal cell (mammalian; erythrocytic cells and method for preserving cells) IT Hydration, physiological (rehydration; erythrocytic cells and method for preserving cells) IT Mesenchyme (stem cell; erythrocytic cells and method for preserving cells) IT 99-20-7, Trehalose 7732-18-5, Water, biological RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (erythrocytic cells and method for preserving cells) 9002-04-4, Thrombin ΙT RL: NUU (Other use, unclassified); USES (Uses) (erythrocytic cells and method for preserving cells) 57-88-5, Cholesterol, properties ΙT RL: PRP (Properties); REM (Removal or disposal); PROC (Process) (erythrocytic cells and method for preserving cells) IT. 99-20-7, Trehalose RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (erythrocytic cells and method for preserving cells) 99-20-7 HCAPLUS RN.alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME) CN Absolute stereochemistry. Rotation (+).

US 2002-52162

Α

20020116

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L73
     ANSWER 6 OF 11
                     HCAPLUS
                              COPYRIGHT 2003 ACS on STN
ΑN
     2002:466549
                 HCAPLUS
DN
     137:17442
     Eukaryotic cells and method for preserving cells
TΙ
     Crowe, John H.; Tablin, Fern; Wolkers, Willem F.; Oliver, Ann E.; Walker,
IN
     Naomi J.; Htoo, Thurein; Jamil, Kamran
PA
SO
     U.S. Pat. Appl. Publ., 36 pp., Cont.-in-part of U.S. Ser. No. 828,627.
     CODEN: USXXCO
DT
     Patent
LA
     English
IC
     ICM A61K035-14
     ICS A01N001-02
NCL
     424532000
     9-11 (Biochemical Methods)
     Section cross-reference(s): 13
FAN.CNT 3
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                                                             DATE
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                            _____
     US 2002076445
                       A1
                            20020620
                                           US 2001-927760
                                                             20010809
PI
                                           US 2001-828627
     US 2001019819
                            20010906
                                                             20010405
                       A1
     US 2002009500
                            20020124
                                           US 2001-938408
                                                             20010823
                       Α1
     US 2002114791
                       A1
                            20020822
                                           US 2002-52162
                                                             20020116
     WO 2003014305
                       A2
                            20030220
                                           WO 2002-US24772
                                                            20020805
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             VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
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             PT, SE,
             NE, SN,
                     TD, TG
                            20030220
                                           WO 2002-US24773 20020805
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                       Α1
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             RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ,
             VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
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             CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT; LU, MC, NL,
             PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
             NE, SN,
                     TD, TG
PRAI US 2000-501773
                            20000210
                       В1
     US 2001-828627
                            20010405
                       Α2
     US 2001-927760
                       A2
                            20010809
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A dehydrated compn. is provided that includes freeze-dried eukaryotic
     cells. The eukaryotic cells are loaded with an oligosaccharide (e.g.,
     trehalose) which preserves biol. properties during freeze-drying
     and rehydration. The oligosaccharide loading is conducted at a temp. of
     from greater than about 25.degree.. to less than about 50.degree.., more
     preferably at about 35.degree.., with the loading soln. having the
     oligosaccharide in an amt. from about 10 mM to about 100 mM. These
     freeze-dried eukaryotic cells are rehydratable. A process for preserving
     and/or increasing the survival of dehydrated eukaryotic cells, including
     storing dehydrated eukaryotic cells having a residual water content
     greater than about 0.15 g of water per g of dry wt. eukaryotic cells.
ST
     eukaryotic cell preserving
IT
    Animal cell line
        (293H; eukaryotic cells and method for preserving cells)
IT
     Hydration, physiological
        (Prehydration; eukaryotic cells and method for preserving cells)
IT
     Temperature effects, biological
        (cold; eukaryotic cells and method for preserving cells)
IT
     Animal cell
     Animal tissue culture
     Cell proliferation
     Composition
      Dehydration, physiological
     Eukaryota
      Freeze drying
     Freezing point
    Mammalia
     Phase transition temperature
     Platelet (blood)
      Preservation
      Preservatives
     Solutions
     Storage
     Temperature effects, biological
     Washing
        (eukaryotic cells and method for preserving cells)
     Oligosaccharides, biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (eukaryotic cells and method for preserving cells)
TT .
    Endocytosis
        (fluid phase; eukaryotic cells and method for preserving cells)
    Hydration, physiological
IT
        (rehydration; eukaryotic cells and method for preserving cells)
ΙT
    Mesenchyme
        (stem cell; eukaryotic cells and method for preserving cells)
ΙT
     Biological transport
        (uptake; eukaryotic cells and method for preserving cells)
ΙT
     7732-18-5, Water, biological studies
                                            9002-04-4, Thrombin
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (eukaryotic cells and method for preserving cells)
IT
     99-20-7, Trehalose
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (eukaryotic cells and method for preserving cells)
     99-20-7, Trehalose
TΨ
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (eukaryotic cells and method for preserving cells)
RN
     99-20-7 HCAPLUS
     .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)
CN
```

Absolute stereochemistry. Rotation (+).

ST

ΙT

Drying

preservation storage medium biol

```
L73
     ANSWER 7 OF 11 HCAPLUS COPYRIGHT 2003 ACS on STN
ΑN
     2002:107034
                 HCAPLUS
     136:131240
DN
ΤI
     Preservation and storage medium for biological materials
     Depablo, Juan J.; Miller, Danforth P.; Conrad, Paul B.; Corti, Horatio
IN
PA
     Wisconsin Alumni Research Foundation, USA
SO
     PCT Int. Appl., 50 pp.
     CODEN: PIXXD2
DT
     Patent
     English
LA
IC
     ICM A01N001-02
CC
     9-11 (Biochemical Methods)
FAN.CNT 1
     PATENT NO.
                      KIND
                            DATE
                                           APPLICATION NO.
                                                             DATE
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                            ______
                                           _____
                                           WO 2001-US14939
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                       Α1
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                                                             20010509
PT
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                       В1
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         RW: GH, GM,
                     KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
                     ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             DE, DK,
                     CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
             BJ, CF,
     EP 1303184
                                                             20010509
                       A1 ·
                            20030423
                                           EP 2001-945951
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
     BR 2001012638
                            20030624
                                           BR 2001-12638
                                                             20010509
                       Α
PRAI US 2000-625735
                       Α
                            20000726
     WO 2001-US14939
                       W
                            20010509
     A protectant mixt. for use in preserving biol. material comprising (1) at
AΒ
     least one polyhydroxy compd., where the total amt. of polyhydroxy compd.
     in the mixt. is from about 5 to about 60 by wt. of the mixt. where the
     mixt. is an aq. soln. and is from about 10 to about 95 where the mixt. is
     in solid form, and (2) phosphate ions, where the total amt. of phosphate
     ions in the mixt. is such that the molar ratio of phosphate ions to
     hydroxy groups in the polyhydroxy compd. is from about 0.025 to about
     0.625; a preservation medium comprising (1) a biol. material, (2) at least
     one polyhydroxy compd., where the total amt. of polyhydroxy compd. in the
     medium is from about 5 to about 60 by wt. of the medium, and (3) phosphate
     ions, where the total amt. of phosphate ions in the mixt. is such that the
     molar ratio of phosphate ions to hydroxy groups in the polyhydroxy compds.
     is from about 0.025 to about 0.625; methods of preserving the preservation
     medium; and the resulting preserved biol. material compn.
```

(Ambient-air; preservation and storage medium for biol. materials)

```
ΙT
     Hydroxy compounds
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (polyhydroxy compds.; preservation and storage medium for biol.
        materials)
     Biological materials
IT
     Cell
     Composition
     Cryoprotectants
     Culture media
       Freeze drying
     Freezing
     Hydroxyl group
     Mixtures
       Preservation.
     Solids
     Solutions
     Storage
     рΗ
        (preservation and storage medium for biol. materials)
IT
     Disaccharides
     Monosaccharides
     Polysaccharides, biological studies
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (preservation and storage medium for biol. materials)
IT
     Enzymes, processes
     Proteins
     RL: CPS (Chemical process); PEP (Physical, engineering or chemical
     process); PROC (Process)
        (preservation and storage medium for biol. materials)
ΙŤ
    Drying
        (spray; preservation and storage medium for biol. materials)
ΙT
     Drying
        (vacuum; preservation and storage medium for biol. materials)
     50-21-5, Lactic acid, biological studies 50-81-7, Ascorbic acid,
IT
     biological studies
                          57-50-1, Sucrose, biological studies
     Citric acid, biological studies 99-20-7, Trehalose
                                                       994-36-5, Sodium citrate
     497-19-8, Sodium carbonate, biological studies
     7558-79-4
                 7558-80-7
                             7757-82-6, Sodium sulfate, biological studies
                                                 7778-77-0
                 7772-98-7, Sodium thiosulfate
                                                              11129-12-7, Borate
                                                 16068-46-5, Potassium
     14265-44-2, Phosphate, biological studies
     phosphate
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (preservation and storage medium for biol. materials)
     9001-60-9, L-Lactate dehydrogenase
     RL: CPS (Chemical process); PEP (Physical, engineering or chemical
     process); PROC (Process)
        (preservation and storage medium for biol. materials)
              THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
RE
(1) Ciba Geigy Ag; WO 9520399 A 1995
(2) Cryopharm Corp; EP 0356257 A 1990 HCAPLUS
(3) Cryopharm Corp; EP 0508496 A 1992 HCAPLUS
(4) Kane, O; US 4476221 A 1984 HCAPLUS
(5) Morishita Pharma; EP 0580444 A 1994 HCAPLUS
     99-20-7, Trehalose
IT
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (preservation and storage medium for biol. materials)
     99-20-7 HCAPLUS
RN
     .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI)
                                                                 (CA INDEX NAME)
```

CN

Absolute stereochemistry. Rotation (+).

IT

Skin

```
L73 ANSWER 8 OF 11 HCAPLUS COPYRIGHT 2003 ACS on STN
     2001:850862 HCAPLUS
DN
     135:368939
ΤI
     Microinjection of biological tissue with intracellular cryoprotective
     agents that contain sugar
ΙN
     Toner, Mehmet; Eroglu, Ali; Toth, Thomas
     The General Hospital Corp., USA; Gamete Technologies, Inc.
PΑ
SO
     PCT Int. Appl., 54 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
IC
     ICM A01N
CC
     9-11 (Biochemical Methods)
FAN.CNT 2
                                                            DATE
     PATENT NO.
                      KIND
                            DATE
                                          · APPLICATION NO.
                      ____
                            _____
                                           ______
                                           WO 2001-US15748
PΙ
     WO 2001087062
                       A2
                            20011122
                                                            20010516
     WO 2001087062
                      А3
                            20020328
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             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT,
             RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ,
             VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
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             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                           US 2001-798327
                                                            20010302
     US 2002045156
                      Α1
                            20020418
                            20011126
                                           AU 2001-61644
                                                            20010516
     AU 2001061644
                       Α5
                                           EP 2001-935562
                                                            20010516
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     EP 1283670
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            AT; BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
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PRAI US 2000-204877P
                     Ρ
                            20000516
     US 2001-798327
                            20010302
                       Α
     WO 2001-US15748
                      W
                            20010516
     The invention concerns the preservation of biol. tissue using
AΒ
     microinjection of intracellular protective agents contg. sugar to preserve
     cells by freezing and/or drying. A method for biol. material including
     micro-injecting the cells with sugar; prepg. the cells for storage;
     storing the biol. material; and recovering the stored biol. material from
     storage. Carbohydrate sugars such as trehalose, sucrose,
     fructose, dextran, and raffinose, may be used as bio-protective agents.
     micro injection cryoprotectant preservation cell membrane carbohydrate
ST
     cytoplasm
```

(epidermis; microinjection of biol. tissue with intracellular

cryoprotective agents that contain sugar)

```
ΙT
    Skin
        (keratinocyte; microinjection of biol. tissue with intracellular
        cryoprotective agents that contain sugar)
IT
    Animal cell
    B cell (lymphocyte)
    Cell membrane
    Chondrocyte
      Cryopreservation
    Cryoprotectants
    Culture media
    Cytoplasm
    Epithelium
    Erythrocyte
    Fibroblast
      Freeze drying
    Glass transition temperature
    Hematopoietic precursor cell
    Macrophage
    Melanocyte
    Melting
    Molecular weight
    Monocyte
    Osmolarity
    T cell (lymphocyte)
    Vacuum
        (microinjection of biol. tissue with intracellular cryoprotective
        agents that contain sugar)
ΙT
    Carbohydrates, biological studies
    Glycolipids
    Glycoproteins, general, biological studies.
    RL: BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological
    study); USES (Uses)
        (microinjection of biol. tissue with intracellular cryoprotective
        agents that contain sugar)
ΙT
    Muscle
        (myogenic cell; microinjection of biol. tissue with intracellular
        cryoprotective agents that contain sugar)
ΙT
        (neuron; microinjection of biol. tissue with intracellular
        cryoprotective agents that contain sugar)
IT
        (oocyte; microinjection of biol. tissue with intracellular
        cryoprotective agents that contain sugar)
IT
    Hydration, physiological
        (re-; microinjection of biol. tissue with intracellular cryoprotective
        agents that contain sugar)
ΙT
    Embryo, animal
        (stem cell; microinjection of biol. tissue with intracellular
        cryoprotective agents that contain sugar)
IT
        (stem, adult; microinjection of biol. tissue with intracellular
        cryoprotective agents that contain sugar)
     50-70-4, Sorbitol, biological studies
                                             50-99-7, Glucose, biological
    studies 57-48-7, D-Fructose, biological studies
                                                        57-50-1, Sucrose,
    biological studies
                          63-42-3, Lactose
                                             69-65-8, Mannitol
                                                                 69-79-4,
    Maltose 99-20-7, Trehalose
                                  470-55-3, Stachyose
     512-69-6, Raffinose
                           9004-54-0, Dextran, biological studies
    RL: BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological
     study); USES (Uses)
        (microinjection of biol. tissue with intracellular cryoprotective
        agents that contain sugar)
IT
     99-20-7, Trehalose
     RL: BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological
```

study); USES (Uses)

(microinjection of biol. tissue with intracellular cryoprotective agents that contain sugar)

RN 99-20-7 HCAPLUS

CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

L73 ANSWER 9 OF 11 HCAPLUS COPYRIGHT 2003 ACS on STN

AN 2001:25421 HCAPLUS

DN 134:204612

TI Properties of Human Free Apolipoprotein(a) and Lipoprotein(a) after Either Freezing or Lyophilization in the Presence and Absence of Cryopreservatives

AU Edelstein, Celina; Hinman, Janet; Marcovina, Santica; Scanu, Angelo M.

CS Department of Medicine, University of Chicago, Chicago, IL, 60637, USA

SO Analytical Biochemistry (2001), 288(2), 201-208 CODEN: ANBCA2; ISSN: 0003-2697

PB Academic Press

DT Journal

LA English

CC 9-11 (Biochemical Methods)
 Section cross-reference(s): 6

AΒ Apolipoprotein(a), apo(a), the specific multikringle glycoprotein constituent of lipoprotein(a), Lp(a), occurs in the plasma mostly bound to apoB100-contg. lipoproteins but also in a free form. Often the properties of these products are detd. after storage in the cold; yet limited information is available on their stability at low temps. To shed light on this subject, we examd. the effect of two parameters, freezing and lyophilization, in either the absence or the presence of cryopreservatives. Lp(a)s each having a single apo(a) size isoform contg. either 14 or 17 kringle (K) IVs were isolated from the plasma of healthy donors by combining d. gradient ultracentrifugation and lysine-Sepharose column chromatog. using solns. contg. both antioxidants and proteolytic inhibitors. Apo(a) was obtained from parent Lp(a) by a mild limited reductive procedure. Either freezing at -20.degree. or lyophilization in the presence of 5% sucrose did not change the electrophoretic, immunochem., and lysine-binding properties of Lp(a) including its ability to generate free apo(a). Irresp. of source, apo(a) remained stable when either frozen at -20 and -80.degree. or lyophilized in the presence of 125 mM trehalose. In all cases, the absence of cryopreservatives caused the samples to aggregate irreversibly. Thawed or reconstituted samples of both free and bound apo(a) kept at 4.degree. under sterile conditions in the presence of antioxidants, proteolytic inhibitors, and cryopreservative exhibited no significant changes in properties within the time of observation. Both apo(a) isoforms gave comparable results. We conclude that apo(a), either free or bound, can be kept stable at low temps. in the presence of appropriate cryopreservatives. (c) 2001 Academic Press.

ST apolipoprotein lipoprotein a Lpa freezing lyophilization storage

```
cryopreservation human
ΙT
     Apolipoproteins
     Lipoproteins
     RL: ANT (Analyte); BSU (Biological study, unclassified); PEP (Physical,
     engineering or chemical process); ANST (Analytical study); BIOL
      (Biological study); PROC (Process)
         (Lp(a); properties of human free apolipoprotein(a) and lipoprotein(a)
        after either freezing or lyophilization in presence and absence of
        cryopreservatives)
ΙT
     Cryopreservation
       Freeze drying
     Freezing
       Preservatives
     Storage
         (properties of human free apolipoprotein(a) and lipoprotein(a) after
        either freezing or lyophilization in presence and absence of
        cryopreservatives)
     57-50-1, Sucrose, analysis 99-20-7, Trehalose
\mathbf{I} \cdot \mathbf{T}
     RL: ARU (Analytical role, unclassified); BUU (Biological use,
     unclassified); ANST (Analytical study); BIOL (Biological study); USES
         (properties of human free apolipoprotein(a) and lipoprotein(a) after
        either freezing or lyophilization in presence and absence of
        cryopreservatives)
              THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
RE
(1) Draber, P; J Immunol Methods 1995, V181, P37 HCAPLUS
(2) Edelstein, C; Biochemistry 1995, V34, P16483 HCAPLUS
(3) Edelstein, C; J Biol Chem 1999, V274, P10019 HCAPLUS
(4) Edelstein, C; J Biol chem 1997, V272, P11079 HCAPLUS
(5) Edelstein, C; J Lipid Res 1996, V37, P1786 HCAPLUS
(6) Fless, G; Biochemistry 1994, V33, P13492 HCAPLUS
(7) Fless, G; J Lipid Res 1989, V30, P651 HCAPLUS
(8) Klezovitch, O; J Biol Chem 1998, V273, P23856 HCAPLUS
(9) Kostner, G; J Lipid Res 1999, V40, P2255 HCAPLUS
(10) Lackner, C; J Clin Invest 1991, V87, P2153 HCAPLUS
(11) Leslie, S; Appl Environ Microbiol 1995, V61, P3592 HCAPLUS
(12) Marcovina, S; Clin Chem 1995, V41, P246 HCAPLUS
(13) Phillips, M; Biochemistry 1993, V32, P3722 HCAPLUS
(14) Sattler, W; Biochim Biophys Acta 1991, V1081, P65 HCAPLUS
(15) Schumaker, V; Methods Enzymol 1986, V128, P155 HCAPLUS
(16) Shoukry, M; Biochim Biophys Acta 1994, V1210, P355 HCAPLUS
(17) Sola-Penna, M; Z Naturforsch 1994, V49c, P327
(18) Uritani, M; J Biochem 1995, V117, P774 HCAPLUS
(19) Wilkie, S; BioChromatography 1988, V5, P205
ΙΤ
     99-20-7, Trehalose
     RL: ARU (Analytical role, unclassified); BUU (Biological use,
     unclassified); ANST (Analytical study); BIOL (Biological study); USES
         (properties of human free apolipoprotein(a) and lipoprotein(a) after
        either freezing or lyophilization in presence and absence of
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RN
     99-20-7 HCAPLUS
      .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)
CN
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Absolute stereochemistry. Rotation (+).

L73 ANSWER 10 OF 11 HCAPLUS COPYRIGHT 2003 ACS on STN

AN 2000:693284 HCAPLUS

DN 134:249126

TI Stabilization and Preservation of Lactobacillus acidophilus in Saccharide Matrices

AU Conrad, Paul B.; Miller, Danforth P.; Cielenski, Peter R.; de Pablo, Juan J.

CS Department of Chemical Engineering, University of Wisconsin-Madison, Madison, WI, 53706, USA

SO Cryobiology (2000), 41(1), 17-24 CODEN: CRYBAS; ISSN: 0011-2240

PB Academic Press

DT Journal

LA English

CC 9-11 (Biochemical Methods)
Section cross-reference(s): 10

Lyophilization and vacuum- or spray-drying are some of the most useful ÀΒ techniques for preserving foods, agricultural products, and pharmaceuticals. Biol. materials, however, can be irreversibly damaged during these treatments. Therefore, it is essential to design protective agents to preserve protein activity and cell viability. In this paper we examine the use of .alpha.,.alpha.-trehalose-borate systems as protectants for Lactobacillus acidophilus during freeze- and vacuum-drying. Trehalose was found to be an effective protectant for freeze-dried and vacuum-dried samples, and it is equiv. to a protective formulation which is in current industrial use. It is known from our previous work on enzymes that the presence of borate can dramatically enhance the protective ability of trehalose. In this work, the addn. of trehalose-borate to bacterial conc. greatly improves the recovery of viable cells after storage. improvement was seen in freeze-dried samples stored at 37 degree. as well as for vacuum-dried samples held at room temp. A tailored buffering strategy was tested to counteract the high pH resulting from the addn. of borate to the mixt. Use of citric or lactic acids in combination with ammonium hydroxide gave a protectant soln. with high pH (resulting in effective crosslinking between trehalose and borate) but a dry product with reduced pH upon rehydration (conducive to cell survival). These results raise exciting possibilities for protection of more labile prokaryotic species as well as simple eukaryotes. (c) 2000 Academic Press.

ST stabilization preservation Lactobacillus acidophilus saccharide

IT Hydration, physiological

(rehydration; stabilization and preservation of Lactobacillus acidophilus in saccharide matrixes)

IT Drying

(spray; stabilization and preservation of Lactobacillus acidophilus in saccharide matrixes)

IT Biological materials Buffers

```
Concentration (condition)
    Crosslinking
    Eukaryote (Eukaryotae)
      Freeze drying
    Lactobacillus acidophilus
      Preservation
    Prokaryote
    Stabilizing agents
    Storage
    рН
        (stabilization and preservation of Lactobacillus acidophilus in
        saccharide matrixes)
ΙT
    Carbohydrates, biological studies
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (stabilization and preservation of Lactobacillus acidophilus in
        saccharide matrixes)
IT
    Drying
        (vacuum; stabilization and preservation of Lactobacillus acidophilus in
        saccharide matrixes)
                                                77-92-9, Citric acid,
    50-21-5, Lactic acid, biological studies
    biological studies 99-20-7, Trehalose
                                             1336-21-6,
    Ammonium hydroxide
                        11129-12-7, Borate
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (stabilization and preservation of Lactobacillus acidophilus in
        saccharide matrixes)
RE.CNT
             THERE ARE 18 CITED REFERENCES AVAILABLE FOR THIS RECORD
(1) Aldous, B; Cryo-Lett 1995, V16, P181 HCAPLUS
(2) Benhra, A; Cryò-Lett 1994, V15, P269
(3) Chesne, C; Cryobiology 1988, V25, P323 MEDLINE
(4) Conrad, P; J Phys Chem 1999, V103, P4049 HCAPLUS
(5) Crowe, J; Biochim Biophys Acta 1988, V947, P367 HCAPLUS
(6) Crowe, J; Biopharm 1993, V6, P28 HCAPLUS
(7) Crowe, J; Biopharm 1993, V6, P40 HCAPLUS
(8) Crowe, L; Biophys J 1996, V71, P2087 HCAPLUS
(9) Damjanovic, V; Cryobiology 1972, V9, P565 MEDLINE
(10) Fuller, B; Cryo-Lett 1980, V1, P139 HCAPLUS
(11) Green, J; J Phys Chem 1989, V93, P2880 HCAPLUS
(12) Hoekstra, F; Comp Biochem Physiol A 1997, V117, P335
(13) Mazur, P; Science 1970, V168, P939 MEDLINE
(14) Miller, D; Pharm Res 1997, V14, P578 HCAPLUS
(15) Miller, D; Pharm Res 1998, V15, P1215 HCAPLUS
(16) Pringle, M; Effects of Low Temperature on Biological Systems 1981, P21
   HCAPLUS
(17) Rudolph, A; Arch Biochem Biophys 1986, V245, P134 HCAPLUS
(18) Stiff, P; Blood 1987, V70, P974 MEDLINE
    99-20-7, Trehalose
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (stabilization and preservation of Lactobacillus acidophilus in
        saccharide matrixes)
     99-20-7 HCAPLUS
RN
     .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)
CN
Absolute stereochemistry. Rotation (+).
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L73 ANSWER 11 OF 11 HCAPLUS COPYRIGHT 2003 ACS on STN

AN 2000:107596 HCAPLUS

DN 132:233904

TI Trehalose expression confers desiccation tolerance on human cells

AU Guo, Ning; Puhlev, Iskren; Brown, David R.; Mansbridge, Jonathan; Levine, Fred

CS Center for Molecular Genetics, UCSD School of Medicine, La Jolla, CA, 92093-0634, USA

SO Nature Biotechnology (2000), 18(2), 168-171 CODEN: NABIF9; ISSN: 1087-0156

PB Nature America

DT Journal

LA English

CC 9-11 (Biochemical Methods)
Section cross-reference(s): 13

AB Many organisms that withstand desiccation express the disaccharide trehalose. We have now expressed the otsA and otsB genes of Escherichia coli, which encode trehalose biosynthetic enzymes, in human primary fibroblasts using a recombinant adenovirus vector. Infected cells produced increased amts. of trehalose with increasing multiplicity of infection (MOI). Human primary fibroblasts expressing trehalose could be maintained in the dry state for up to five days. Fourier transform IR spectroscopy indicated that dry, but viable, human cells contained no detectable water. This study shows that mammalian cells can be engineered to retain viability in the absence of water.

ST trehalose desiccation tolerance dehydration tissue preservation fibroblast

IT Dehydration, physiological

Fibroblast

(trehalose expression and desiccation tolerance on human cells)

IT Organ preservation

(trehalose expression and desiccation tolerance on human cells in relation to)

IT 99-20-7, Trehalose

RL: BOC (Biological occurrence); BSU (Biological study, unclassified); MFM (Metabolic formation); BIOL (Biological study); FORM (Formation, nonpreparative); OCCU (Occurrence)

(trehalose expression and desiccation tolerance on human cells)

RE.CNT 21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

- (1) Allison, S; Arch Biochem Biophys 1999, V365, P289 HCAPLUS
- (2) Arakawa, T; Cryobiology 1990, V27, P401 HCAPLUS
- (3) Beattie, G; Diabetes 1997, V46, P519 HCAPLUS
- (4) Berkner, K; Curr Top Microbiol Immunol 1991, V158, P39
- (5) Crowe, J; Annu Rev Physiol 1992, V54, P579 HCAPLUS

- (6) Crowe, J; Annu Rev Physiol 1998, V60, P73 HCAPLUS
- (7) Crowe, J; Biochem J 1987, V242, P1 HCAPLUS
- (8) Crowe, L; Dev Biol Standardization 1992, V74, P285 HCAPLUS
- (9) Hirata, T; Surgery 1994, V115, P102 MEDLINE
- (10) Kaasen, J; Gene 1994, V145, P9
- (11) Leibowitz, G; Diabetes 1999, V48, P745 HCAPLUS
- (12) Leslie, S; Appl Environ Microbiol 1995, V61, P3592 HCAPLUS
- (13) Leslie, S; Biochim Biophys Acta 1994, V1192, P7 HCAPLUS
- (14) Naughton, G; Artif Organs 1997, V21, P1203 MEDLINE
- (15) Pelletier, J; Nature 1988, V334, P320 HCAPLUS
- (16) Singer, M; Mol. Cell 1998, V1, P639 HCAPLUS
- (17) Slade, L; Adv Exp Med Biol 1991, V302, P29 HCAPLUS
- (18) Spessot, R; Virology 1989, V168, P378 HCAPLUS
- (19) Sun, W; Biophys J 1996, V70, P1769 HCAPLUS
- (20) Thevelein, J; Trends Biochem Sci 1995, V20, P3 HCAPLUS
- (21) Wiemken, A; Antonie Van Leeuwenhoek 1990, V58, P209 HCAPLUS
- IT 99-20-7, Trehalose

RL: BOC (Biological occurrence); BSU (Biological study, unclassified); MFM (Metabolic formation); BIOL (Biological study); FORM (Formation, nonpreparative); OCCU (Occurrence)

(trehalose expression and desiccation tolerance on human cells)

RN 99-20-7 HCAPLUS

CN .alpha.-D-Glucopyranoside, .alpha.-D-glucopyranosyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

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MOST RECENT DERWENT UPDATE: 200364 <200364/DW>
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http://thomsonderwent.com/coverage/latestupdates/ <<< >>> FOR INFORMATION ON ALL DERWENT WORLD PATENTS INDEX USER GUIDES, PLEASE VISIT: http://thomsonderwent.com/support/userguides/ <<< => d 190 all tot abeq tech abex ANSWER 1 OF 2 WPIX COPYRIGHT 2003 THOMSON DERWENT on STN 2001-080827 [09] WPIX ΑN DNC C2001-023329 TΙ Preserving biologically active material, particularly viruses such as measles, mumps, and rubella, comprises mixing a biological suspension with a sterile mixture of chitosan. DC B04 C06 D16 D22 ΙN WORRALL, E E PA (WORR-I) WORRALL E E CYC WO 2000078924 A1 20001228 (200109)* EN 24p C12N001-04 PΤ RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TZ UG ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW AU 2000054135 A 20010109 (200122) C12N001-04 <--EP 1187907 A1 20020320 (200227) EN C12N001-04 <--R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI CN 1357037 A 20020703 (200265) C12N001-04 ADT WO 2000078924 A1 WO 2000-GB2254 20000621; AU 2000054135 A AU 2000-54135 20000621; EP 1187907 A1 EP 2000-938911 20000621, WO 2000-GB2254 20000621; CN 1357037 A CN 2000-809328 20000621 FDT AU 2000054135 A Based on WO 2000078924; EP 1187907 Al Based on WO 2000078924 19990622 PRAI GB 1999-14412 ICM C12N001-04 IC AB WO 200078924 A UPAB: 20010213 NOVELTY - Preserving biologically-active material, comprising mixing an aqueous suspension of the material with a sterile aqueous solution of chitosan to form a coacervate, adding a sterile aqueous solution of trehalose, drying the mixture at low pressure, and at a temperature, initially no more than 37 deg. C, which is subsequently controlled not to fall to 0 deg. C or below to form a glassy porous matrix, is new. DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following: (1) making a vaccine, comprising preserving a biologically active material using the novel method, and rehydrating the glassy product in an aqueous medium; and (2) a rehydratable composition comprising trehalose in the form of a metastable containing, within the matrix, desiccated biologically material and chitosan or its non-toxic salt. USE - The process is used to preserve viruses (e.g. Rinderpest virus, Peste de Petit Ruminants virus, Measles, Mumps, Rubella, Yellow Fever, Polio and Newcastle Disease Virus), bacteria, Contagious Bovine Pleuropneumonia (CBPP) mycoplasma, tertiary structured biologically-active protein and nucleic acid (all claimed). Dwg.0/0 FŞ CPI

CPI: B04-C02E3; B04-E01; B04-F11; B04-N05; B07-A02B; B14-S11;

FΑ

MC

AB; DCN

C04-C02E3; C04-E01; C04-F11; C04-N04; C07-A02B; C14-S11; D05-H07; D09-A01

UPTX: 20010213

TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred Materials: The sterile aqueous solution of chitosan or its non-toxic salt has a chitosan concentration of 0.01 % w/v. The sterile aqueous chitosan solution and the aqueous suspension of biologically-active material are mixed at a volume ratio of 1:1 at pH 7.4. The coacervate of biologically-active material and chitosan is subjected to vortex mixing. The coacervate is mixed with a sterile aqueous trehalose solution having a trehalose concentration in the range of from 0.20-20, especially 5 % w/v. The mixture is subjected to drying for 30-60 minutes, at a pressure of less than atmospheric and at no more than 37 degrees C, and which is controlled not to fall to 0 degrees C, and which is finally no greater than 40 degrees C, to form a glassy porous matrix comprising glassy trehalose having a residual moisture content not greater than 10 % and containing, within the matrix, desiccated biologically-active material and chitosan or its non-toxic salt. The drying stage is carried out at a pressure of not greater than 800 mbar. The resulting trehalose matrix is subjected to a secondary drying procedure. The residual moisture content at the end of the secondary drying step is 1.0 % or lower. The vaccine is for oral or intranasal use. The vaccine is a Measles, Mumps, Rubella (MMR) vaccine.

ABEX UPTX: 20010213

TECH

EXAMPLE - Rinderpest virus RBOK strain was grown in vero cells in Hanks LYE (lactalbumin hydrolysate and yeast extract) medium containing 0.1 % trehalose instead of glucose. Contagious Bovine Pleuropneumonia (CBPP) Mycoplasma mycoides subs.mycoides S1144 (SC) TI-SR was grown in Gourlay medium. The virus pool and the CBPP pool were then harvested. The pH of the virus and the CBPP pool were adjusted with 0.1 M NaOH to pH 7.4.A stock solution of 50 % w/v trehalose dehydrate in Hanks Balanced Salt Solution (HBSS) was prepared the pH being adjusted by the addition of 0.1 M NaOH to 7.4. The solution was sterilized by autoclaving at 121 degrees C for 20 minutes. A suitable volume of working strength 0.02 % w/v chitosan was prepared by adding 1 ml of stock 2 % w/v chitosan to 99 ml of sterile distilled water. One volume of 0.02 % w/v chitosan solution was added to one volume of virus fluid at 4 degrees C and the pH adjusted to 7.4 with 0.1 M NaOH. This step was repeated separately for the CBPP culture pool and the resulting coacervation complex of each was completed by rapid vortex stirring for 30 seconds and subsequently stored at 4 degrees C for 1 hour. The resulting precipitate of each was collected by centrifugation at 10000 revolutions per minute (rpm) in a refrigerated centrifuge, the supernatant discarded and the coacervate resuspended in one volume of Hanks balanced salt solution (HBSS). A volume of sterile 50 % w/v solution of trehalose in Hanks balanced salt solution (HBSS) was added to the coacervate suspension give a final trehalose concentration of 5 $% \ w/v \ (potency is checked by the quality control standard operating$ procedures for these organisms). The vaccine was dried by filling 1.0 ml aliquots into 5 mi vaccine vials partially stoppered with dry butyl rubber stoppers. The shelves of the conventional freeze dryer (EDWARDS MODULYO) were heated to 37 degrees C and the condenser was allowed to reach minus 40 degrees C. The vaccine vials were placed in the dryer and the pressure in the drying chamber was adjusted to 800 mbar and drying commenced for 30 minutes until 75 % of the water had evaporated, taking care not to allow the product temperature to fall below 0 degrees C. The pressure was then lowered to 500 mbar and drying was continued until the glass transition temperature of trehalose was reached at 25 degrees C, a glassy porous matrix was formed, and the temperature of the product was allowed to rise to reach the initial starting temperature close to that of the shelves. At this stage the residual moisture (RM) was 10 %. Further drying at 0.01 mbar and 45 degrees C for 17 hours reduced this to 1-2 % RM

ensuring high thermostability in the product. The pressure was maintained at 0.01 mbar and the vials were sealed under vacuum or at atmospheric pressure under dry nitrogen.

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L90 ANSWER 2 OF 2 WPIX COPYRIGHT 2003 THOMSON DERWENT on STN
AN
     2000-679756 [66]
                       WPIX
DNC C2000-206828
ΤI
     Preservation of viruses and mycoplasma for vaccines in a trehalose matrix,
     with increased thermostability.
DC
     B04 C06 D16
ΙN
     WORRALL, E E
PA
     (WORR-I) WORRALL E E; (ANHY-N) ANHYDRO LTD
CYC
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    .20000503; EP 1175486 A2 EP 2000-927438 20000503, WO 2000-GB1524 20000503;
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PRAI GB 1999-26698
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    C12N001-04; C12N007-00; C12R001:93; C12R001:93
ICI
AB
     WO 200066710 A UPAB: 20001219
     NOVELTY - Preserving viruses and mycoplasma comprising desiccation without
     lyophilization, is new.
          DETAILED DESCRIPTION - Preserving viruses and mycoplasma comprising
     desiccation without lyophilization, is new. The method comprises:
```

- (1) mixing an aqueous suspension of the biologically-active material with a sterile aqueous solution of trehalose to give a trehalose
- with a sterile aqueous solution of trehalose to give a trehalose concentration of 0.2-10 % w/v;

 (2) subjecting the mixture to primary drying for 30-60 minutes at
- (2) subjecting the mixture to primary drying for 30-60 minutes at 0-37 deg. C initially and 0-40 deg. C finally, at at least atmospheric pressure, to form a glassy porous matrix comprising glassy trehalose with a residual moisture content of no more than 10 %, and containing within the matrix, desiccated biologically active material; and
- (3). subjecting the matrix to secondary drying for 10-30 hours at a no more than 0.1 mbar and 40-45 deg. C, to form a trehalose matrix with residual moisture content of no more than 2 %.

INDEPENDENT CLAIMS are also included for the following:

- (1) a rehydratable composition made by the novel method; and
- (2) making a vaccine comprising rehydrating the composition in an appropriate aqueous medium.
- USE For preserving viruses and mycoplasma for use in vaccines (claimed).

ADVANTAGE - The method is much faster than prior freeze drying methods, a moisture content of less than 10 % can be achieved in less than an hour, and damage caused by solute concentration and ice crystallisation is minimized. The preserved material is thermostable and it can be exposed to temperatures of up to 45 deg. C without loss of biological activity. The material can be instantly rehydrated.

Dwg.0/1
CPI
AB; DCN
CPI: B04-F10A4; B04-F11; B10-A07; B14-S11; C04-F10A4; C04-F11; C10-A07;

C14-S11; D05-H07; D05-H10 TECH UPTX: 20001219

TECHNOLOGY FOCUS - BIOLOGY - Preferred Method: The secondary drying step is for 20-30, preferably 15-17 hours at 37 degrees C, then at 40-45 degrees C for the remaining time. The trehalose concentration is 2-10, especially 2.5-8 % w/v. The primary drying step pressure is no more than 800 mbar, and the residual moisture content after this step is 10 %. Preferred Material: The virus is selected from Rinderpest, Peste des Petits Ruminants, Measles, Mumps, Rubella, Yellow Fever, Polio and Newcastle Disease viruses, especially Rinderpest or Peste des Petits Ruminants. The mycoplasma is Contagious Bovine Pleuropneumonia mycoplasma.

ABEX UPTX: 20001219

EXAMPLE - The thermostability of viruses was measured when subjected to different lengths of secondary dehydration. Rinderpest viruses subjected to 17 hours secondary drying had a virus titer of 4.97 on the first day, and 3.03 after 14 days, when stored at 45 degrees C, compared to 5.40 and 0.0 respectively for peste des petits ruminants virus subjected to only 2 hours secondary drying. This shows how the low moisture levels (less than 1 %) after extended secondary drying gives increased thermostability.

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          297 S L18 AND L38
           139 S L18 AND L39
L41
L42
           68 S L14 AND L38
           20 S L14 AND L39
            8 S L19 AND L31-L34
             5 S L19 AND L40-L43
             12 S L44, L45
L46
L47
             1 S L46 AND L37
             11 S L46 NOT L47
             SEL DN AN 6 7
             2 · S E1-E6 AND L48
L49
             6 S L37, L47, L49
L50
             6 S L50 AND L10-L50
L51
              E DRYING/CT
               E E3+ALL
          32988 S E2
L52
               E E1+ALL
            443 S E1
L53
              E E6+ALL
L54
          19277 S E2, E1+NT
             E E13+ALL
          25489 S E6, E7, E5
L55
           334 S L18 AND L52-L55
L56
            133 S L14 AND L52-L55
L57
            6 S L19 AND L56,L57
L58
             3 S L51 AND L52-L57
L59
             6 S L51,L59
L60
           637 S (BIOCHEM?(L)METHOD?)/SC,SX AND L14
L61
           649 S (BIOCHEM?(L)METHOD?)/SC, SX AND L18
L62
           95 S L61, L62 AND L31-L34
L63
            22 S L63 AND L38-L41
L64
            21 S L63 AND L52-L57
L65
            27 S L64,L65
L66
```

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16 S L66 AND (PY<=1999 OR PRY<=1999 OR AY<=1999)
L67
L68
             0 S L66 AND L19
L69
             11 S L66 NOT L67
             32 S L60, L67, L69
             19 S L70 AND (PY<=1999 OR PRY<=1999 OR AY<=1999)
L72
             21 S L37, L71
             11 S L70 NOT L72
L73
                SEL HIT RN L70
     FILE 'REGISTRY' ENTERED AT 15:58:09 ON 07 OCT 2003
L74
              2 S E1-E2
     FILE 'REGISTRY' ENTERED AT 15:58:19 ON 07 OCT 2003
     FILE 'HCAPLUS' ENTERED AT 15:58:29 ON 07 OCT 2003
     FILE 'WPIX' ENTERED AT 16:00:07 ON 07 OCT 2003
                E WORRAL E/AU
L75
              2 S E15
                E CHITOSAN/DCN
                E E3+ALL
L76
           1505 S E2
            873 S E4
                E TREHALOSE/DCN
                E E3+ALL
L78
            700 S E2
L79
           1670 S (B04-C02E3 OR C04-C02E3)/MC
L80
          11360 S (B10-A07 OR C10-A07)/MC
L81
           553 S C12N001-04/IC, ICM, ICS
L82
            38 S C12N001-04/ICA, ICI
L83
              3 S C12N001:04/ICI
L84
           5351 S CHITOSAN/BIX
L85
           1534 S TREHALOSE/BIX
           5901 S L76, L77, L79, L84
L86
          12712 S L78, L80, L85
L87
            156 S L86 AND L87
L88
              1 S L88 AND L81-L83
L89
L90
              2 S L75, L89.
L91
              9 S L88 AND A61K039/IC, ICM, ICS, ICA, ICI
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FILE 'WPIX' ENTERED AT 16:10:25 ON 07 OCT 2003